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(54) Title: PATHOGEN TOLERANCE GENES

(57) Abstract: The present invention relates to transgenic plants and methods of making transgenic plant using punitive transcription factors that modulate the transgenic plant's susceptibility to disease.

**PATHOGEN TOLERANCE GENES****RELATED APPLICATION INFORMATION**

The present invention claims the benefit from US Provisional Patent Application  
5 Serial Nos. 60/166,228 filed November 17, 1999 and 60/197,899 filed April 17, 2000 and  
"Plant Trait Modification III" filed August 22, 2000.

**FIELD OF THE INVENTION**

This invention relates to the field of plant biology. More particularly, the present invention pertains to compositions and methods for phenotypically modifying a plant.

**10 BACKGROUND OF THE INVENTION**

Transcription factors can modulate gene expression, either increasing or decreasing (inducing or repressing) the rate of transcription. This modulation results in differential levels of gene expression at various developmental stages, in different tissues and cell types, and in response to different exogenous (e.g., environmental) and endogenous 15 stimuli throughout the life cycle of the organism.

Because transcription factors are key controlling elements of biological pathways, altering the expression levels of one or more transcription factors can change entire biological pathways in an organism. For example, manipulation of the levels of selected transcription factors may result in increased expression of economically useful proteins or 20 metabolic chemicals in plants or to improve other agriculturally relevant characteristics. Conversely, blocked or reduced expression of a transcription factor may reduce biosynthesis of unwanted compounds or remove an undesirable trait. Therefore, manipulating transcription factor levels in a plant offers tremendous potential in agricultural biotechnology for modifying a plant's traits.

25 The present invention provides novel transcription factors useful for modifying a plant's phenotype in desirable ways, such as modifying a plant's pathogen tolerance.

**SUMMARY OF THE INVENTION**

In a first aspect, the invention relates to a recombinant polynucleotide 30 comprising a nucleotide sequence selected from the group consisting of: (a) a nucleotide sequence encoding a polypeptide comprising a sequence selected from SEQ ID Nos. 2N, where N=1-29, or a complementary nucleotide sequence thereof; (b) a nucleotide sequence encoding a polypeptide comprising a conservatively substituted variant of a polypeptide of (a); (c) a nucleotide sequence comprising a sequence selected from those of SEQ ID Nos. 2N-35 1, where N=1-29, or a complementary nucleotide sequence thereof; (d) a nucleotide sequence

comprising silent substitutions in a nucleotide sequence of (c); (e) a nucleotide sequence which hybridizes under stringent conditions over substantially the entire length of a nucleotide sequence of one or more of: (a), (b), (c), or (d); (f) a nucleotide sequence comprising at least 15 consecutive nucleotides of a sequence of any of (a)-(e); (g) a nucleotide sequence comprising a subsequence or fragment of any of (a)-(f), which subsequence or fragment encodes a polypeptide having a biological activity that modifies a plant's pathogen tolerance; (h) a nucleotide sequence having at least 31% sequence identity to a nucleotide sequence of any of (a)-(g); (i) a nucleotide sequence having at least 60% identity sequence identity to a nucleotide sequence of any of (a)-(g); (j) a nucleotide sequence which encodes a polypeptide having at least 31% identity sequence identity to a polypeptide of SEQ ID Nos. 2N, where N=1-29; (k) a nucleotide sequence which encodes a polypeptide having at least 60% identity sequence identity to a polypeptide of SEQ ID Nos. 2N, where N=1-29; and (l) a nucleotide sequence which encodes a conserved domain of a polypeptide having at least 65% sequence identity to a conserved domain of a polypeptide of SEQ ID Nos. 2N, where N=1-29. The recombinant polynucleotide may further comprise a constitutive, inducible, or tissue-active promoter operably linked to the nucleotide sequence. The invention also relates to compositions comprising at least two of the above described polynucleotides.

In a second aspect, the invention is an isolated or recombinant polypeptide comprising a subsequence of at least about 15 contiguous amino acids encoded by the recombinant or isolated polynucleotide described above.

In another aspect, the invention is a transgenic plant comprising one or more of the above described recombinant polynucleotides. In yet another aspect, the invention is a plant with altered expression levels of a polynucleotide described above or a plant with altered expression or activity levels of an above described polypeptide. Further, the invention may be a plant lacking a nucleotide sequence encoding a polypeptide comprising a sequence selected from SEQ ID Nos. 2N, where N=1-29.

The plant may be a soybean, wheat, corn, potato, cotton, rice, oilseed rape, sunflower, alfalfa, sugarcane, turf, banana, blackberry, blueberry, strawberry, raspberry, cantaloupe, carrot, cauliflower, coffee, cucumber, eggplant, grapes, honeydew, lettuce, mango, melon, onion, papaya, peas, peppers, pineapple, spinach, squash, sweet corn, tobacco, tomato, watermelon, rosaceous fruits, or vegetable brassicas plant.

In a further aspect, the invention relates to a cloning or expression vector comprising the isolated or recombinant polynucleotide described above or cells comprising the cloning or expression vector.

In yet a further aspect, the invention relates to a composition produced by incubating a polynucleotide of the invention with a nuclease, a restriction enzyme, a polymerase; a polymerase and a primer; a cloning vector, or with a cell.

Furthermore, the invention relates to a method for producing a plant having improved pathogen tolerance. The method comprises altering the expression of an isolated or recombinant polynucleotide of the invention or altering the expression or activity of a polypeptide of the invention in a plant to produce a modified plant, and selecting the 5 modified plant for modified pathogen tolerance.

In another aspect, the invention relates to a method of identifying a factor that is modulated by or interacts with a polypeptide encoded by a polynucleotide of the invention. The method comprises expressing a polypeptide encoded by the polynucleotide in a plant; and identifying at least one factor that is modulated by or interacts with the polypeptide. In one 10 embodiment the method for identifying modulating or interacting factors is by detecting binding by the polypeptide to a promoter sequence, or by detecting interactions between an additional protein and the polypeptide in a yeast two hybrid system, or by detecting expression of a factor by hybridization to a microarray, subtractive hybridization or differential display.

15 In yet another aspect, the invention is a method of identifying a molecule that modulates activity or expression of a polynucleotide or polypeptide of interest. The method comprises placing the molecule in contact with a plant comprising the polynucleotide or polypeptide encoded by the polynucleotide of the invention and monitoring one or more of the expression level of the polynucleotide in the plant, the expression level of the polypeptide 20 in the plant, and modulation of an activity of the polypeptide in the plant.

25 In yet another aspect, the invention relates to an integrated system, computer or computer readable medium comprising one or more character strings corresponding to a polynucleotide of the invention, or to a polypeptide encoded by the polynucleotide. The integrated system, computer or computer readable medium may comprise a link between one or more sequence strings to a modified plant pathogen tolerance phenotype.

30 In yet another aspect, the invention is a method for identifying a sequence similar or homologous to one or more polynucleotides of the invention, or one or more polypeptides encoded by the polynucleotides. The method comprises providing a sequence database; and, querying the sequence database with one or more target sequences corresponding to the one or more polynucleotides or to the one or more polypeptides to identify one or more sequence members of the database that display sequence similarity or 35 homology to one or more of the one or more target sequences.

The method may further comprise of linking the one or more of the polynucleotides of the invention, or encoded polypeptides, to a modified plant pathogen tolerance phenotype.

**BRIEF DESCRIPTION OF THE DRAWINGS**

Figure 1 provides a table of exemplary polynucleotide and polypeptide sequences of the invention. The table includes from left to right for each sequence: the SEQ ID No., the internal code reference number (GID), whether the sequence is a polynucleotide or 5 polypeptide sequence, and identification of any conserved domains for the polypeptide sequences.

Figure 2 provides a table of exemplary sequences that are homologous to other sequences provided in the Sequence Listing and that are derived from *Arabidopsis thaliana*. The table includes from left to right: the SEQ ID No., the internal code reference 10 number (GID), identification of the homologous sequence, whether the sequence is a polynucleotide or polypeptide sequence, and identification of any conserved domains for the polypeptide sequences.

Figure 3 provides a table of exemplary sequences that are homologous to the sequences provided in Figures 1 and 2 and that are derived from plants other than *Arabidopsis thaliana*. The table includes from left to right: the SEQ ID No., the internal code reference 15 number (GID), the unique GenBank sequence ID No. (NID), the probability that the comparison was generated by chance (P-value), and the species from which the homologous gene was identified.

20

**DETAILED DESCRIPTION**

The present invention relates to polynucleotides and polypeptides, e.g. for modifying phenotypes of plants.

In particular, the polynucleotides or polypeptides are useful for modifying traits associated with a plant's pathogen tolerance when the expression levels of the polynucleotides 25 or expression levels or activity levels of the polypeptides are altered. Specifically, the polynucleotides and polypeptides are useful for modifying traits associated with a plant's pathogen tolerance, such as alterations in cell wall composition, trichome number or structure, callose induction, phytoalexin induction, alterations in the cell death response, or the like. Transgenic plants employing the polynucleotides or polypeptides of the invention are more 30 tolerant to biotrophic or necrotrophic pathogens such as fungi, bacteria, mollicutes, viruses, nematodes, parasitic higher plants or the like.

The polynucleotides of the invention encode plant transcription factors. The plant transcription factors are derived, e.g., from *Arabidopsis thaliana* and can belong, e.g., to one or more of the following transcription factor families: the AP2 (APETALA2) domain transcription factor family (Riechmann and Meyerowitz (1998) *J. Biol. Chem.* 379:633-646); the MYB transcription factor family (Martin and Paz-Ares (1997) *Trends Genet.* 13:67-73); the MADS domain transcription factor family (Riechmann and Meyerowitz (1997) *J. Biol.*

Chem. 378:1079-1101); the WRKY protein family (Ishiguro and Nakamura (1994) Mol. Gen. Genet. 244:563-571); the ankyrin-repeat protein family (Zhang et al. (1992) Plant Cell 4:1575-1588); the miscellaneous protein (MISC) family (Kim et al. (1997) Plant J. 11:1237-1251); the zinc finger protein (Z) family (Klug and Schwabe (1995) FASEB J. 9: 597-604);  
5 the homeobox (HB) protein family (Duboule (1994) Guidebook to the Homeobox Genes, Oxford University Press); the CAAT-element binding proteins (Forsburg and Guarente (1989) Genes Dev. 3:1166-1178); the squamosa promoter binding proteins (SPB) (Klein et al. (1996) Mol. Gen. Genet. 1996 250:7-16); the NAM protein family; the IAA/AUX proteins (Rouse et al. (1998) Science 279:1371-1373); the HLH/MYC protein family (Littlewood et  
10 al. (1994) Prot. Profile 1:639-709); the DNA-binding protein (DBP) family (Tucker et al. (1994) EMBO J. 13:2994-3002); the bZIP family of transcription factors (Foster et al. (1994) FASEB J. 8:192-200); the BPF-1 protein (Box P-binding factor) family (da Costa e Silva et al. (1993) Plant J. 4:125-135); and the golden protein (GLD) family (Hall et al. (1998) Plant  
Cell 10:925-936).

15 In addition to methods for modifying a plant phenotype by employing one or more polynucleotides and polypeptides of the invention described herein, the polynucleotides and polypeptides of the invention have a variety of additional uses. These uses include their use in the recombinant production (i.e., expression) of proteins; as regulators of plant gene expression, as diagnostic probes for the presence of complementary or partially  
20 complementary nucleic acids (including for detection of natural coding nucleic acids); as substrates for further reactions, e.g., mutation reactions, PCR reactions, or the like, or as substrates for cloning e.g., including digestion or ligation reactions, and for identifying exogenous or endogenous modulators of the transcription factors.

#### DEFINITIONS

25 A “polynucleotide” is a nucleic acid sequence comprising a plurality of polymerized nucleotide residues, e.g., at least about 15 consecutive polymerized nucleotide residues, optionally at least about 30 consecutive nucleotides, at least about 50 consecutive nucleotides. In many instances, a polynucleotide comprises a nucleotide sequence encoding a polypeptide (or protein) or a domain or fragment thereof. Additionally, the polynucleotide  
30 may comprise a promoter, an intron, an enhancer region, a polyadenylation site, a translation initiation site, 5' or 3' untranslated regions, a reporter gene, a selectable marker, or the like. The polynucleotide can be single stranded or double stranded DNA or RNA. The polynucleotide optionally comprises modified bases or a modified backbone. The polynucleotide can be, e.g., genomic DNA or RNA, a transcript (such as an mRNA), a cDNA,  
35 a PCR product, a cloned DNA, a synthetic DNA or RNA, or the like. The polynucleotide can comprise a sequence in either sense or antisense orientations.

A "recombinant polynucleotide" is a polynucleotide that is not in its native state, e.g., the polynucleotide comprises a nucleotide sequence not found in nature, or the polynucleotide is in a context other than that in which it is naturally found, e.g., separated from nucleotide sequences with which it typically is in proximity in nature, or adjacent (or 5 contiguous with) nucleotide sequences with which it typically is not in proximity. For example, the sequence at issue can be cloned into a vector, or otherwise recombined with one or more additional nucleic acid.

An "isolated polynucleotide" is a polynucleotide whether naturally occurring or recombinant, that is present outside the cell in which it is typically found in nature, whether 10 purified or not. Optionally, an isolated polynucleotide is subject to one or more enrichment or purification procedures, e.g., cell lysis, extraction, centrifugation, precipitation, or the like.

A "recombinant polypeptide" is a polypeptide produced by translation of a recombinant polynucleotide. An "isolated polypeptide," whether a naturally occurring or a recombinant polypeptide, is more enriched in (or out of) a cell than the polypeptide in its 15 natural state in a wild type cell, e.g., more than about 5% enriched, more than about 10% enriched, or more than about 20%, or more than about 50%, or more, enriched, i.e., alternatively denoted: 105%, 110%, 120%, 150% or more, enriched relative to wild type standardized at 100%. Such an enrichment is not the result of a natural response of a wild type plant. Alternatively, or additionally, the isolated polypeptide is separated from other 20 cellular components with which it is typically associated, e.g., by any of the various protein purification methods herein.

The term "transgenic plant" refers to a plant that contains genetic material, not found in a wild type plant of the same species, variety or cultivar. The genetic material may include a transgene, an insertional mutagenesis event (such as by transposon or T-DNA 25 insertional mutagenesis), an activation tagging sequence, a mutated sequence, a homologous recombination event or a sequence modified by chimeroplasty. Typically, the foreign genetic material has been introduced into the plant by human manipulation.

A transgenic plant may contain an expression vector or cassette. The expression cassette typically comprises a polypeptide-encoding sequence operably linked 30 (i.e., under regulatory control of) to appropriate inducible or constitutive regulatory sequences that allow for the expression of polypeptide. The expression cassette can be introduced into a plant by transformation or by breeding after transformation of a parent plant. A plant refers to a whole plant as well as to a plant part, such as seed, fruit, leaf, or root, plant tissue, plant cells or any other plant material, e.g., a plant explant, as well as to progeny thereof, and to *in vitro* 35 systems that mimic biochemical or cellular components or processes in a cell.

The phrase "ectopically expression or altered expression" in reference to a polynucleotide indicates that the pattern of expression in, e.g., a transgenic plant or plant

tissue, is different from the expression pattern in a wild type plant or a reference plant of the same species. For example, the polynucleotide or polypeptide is expressed in a cell or tissue type other than a cell or tissue type in which the sequence is expressed in the wild type plant, or by expression at a time other than at the time the sequence is expressed in the wild type plant, or by a response to different inducible agents, such as hormones or environmental signals, or at different expression levels (either higher or lower) compared with those found in a wild type plant. The term also refers to altered expression patterns that are produced by lowering the levels of expression to below the detection level or completely abolishing expression. The resulting expression pattern can be transient or stable, constitutive or inducible. In reference to a polypeptide, the term "ectopic expression or altered expression" further may relate to altered activity levels resulting from the interactions of the polypeptides with exogenous or endogenous modulators or from interactions with factors or as a result of the chemical modification of the polypeptides.

The term "fragment" or "domain," with respect to a polypeptide, refers to a subsequence of the polypeptide. In some cases, the fragment or domain, is a subsequence of the polypeptide which performs at least one biological function of the intact polypeptide in substantially the same manner, or to a similar extent, as does the intact polypeptide. For example, a polypeptide fragment can comprise a recognizable structural motif or functional domain such as a DNA binding domain that binds to a DNA promoter region, an activation domain or a domain for protein-protein interactions. Fragments can vary in size from as few as 6 amino acids to the full length of the intact polypeptide, but are preferably at least about 30 amino acids in length and more preferably at least about 60 amino acids in length. In reference to a nucleotide sequence, "a fragment" refers to any subsequence of a polynucleotide, typically, of at least consecutive about 15 nucleotides, preferably at least about 30 nucleotides, more preferably at least about 50, of any of the sequences provided herein.

The term "trait" refers to a physiological, morphological, biochemical or physical characteristic of a plant or particular plant material or cell. In some instances, this characteristic is visible to the human eye, such as seed or plant size, or can be measured by available biochemical techniques, such as the protein, starch or oil content of seed or leaves or by the observation of the expression level of genes, e.g., by employing Northern analysis, RT-PCR, microarray gene expression assays or reporter gene expression systems, or by agricultural observations such as stress tolerance, yield or pathogen tolerance.

"Trait modification" refers to a detectable difference in a characteristic in a plant ectopically expressing a polynucleotide or polypeptide of the present invention relative to a plant not doing so, such as a wild type plant. In some cases, the trait modification can be evaluated quantitatively. For example, the trait modification can entail at least about a 2%

increase or decrease in an observed trait (difference), at least a 5% difference, at least about a 10% difference, at least about a 20% difference, at least about a 30%, at least about a 50%, at least about a 70%, or at least about a 100%, or an even greater difference. It is known that there can be a natural variation in the modified trait. Therefore, the trait modification  
5 observed entails a change of the normal distribution of the trait in the plants compared with the distribution observed in wild type plant.

Trait modifications of particular interest include those to seed ( such as embryo or endosperm), fruit, root, flower, leaf, stem, shoot, seedling or the like, including:  
10 enhanced tolerance to environmental conditions including freezing, chilling, heat, drought, water saturation, radiation and ozone; improved tolerance to microbial, fungal or viral diseases; improved tolerance to pest infestations, including nematodes, mollicutes, parasitic higher plants or the like; decreased herbicide sensitivity; improved tolerance of heavy metals or enhanced ability to take up heavy metals; improved growth under poor photoconditions (e.g., low light and/or short day length), or changes in expression levels of genes of interest.  
15 Other phenotype that can be modified relate to the production of plant metabolites, such as variations in the production of taxol, tocopherol, tocotrienol, sterols, phytosterols, vitamins, wax monomers, anti-oxidants, amino acids, lignins, cellulose, tannins, prenyllipids (such as chlorophylls and carotenoids), glucosinolates, and terpenoids, enhanced or compositionally altered protein or oil production (especially in seeds), or modified sugar (insoluble or soluble)  
20 and/or starch composition. Physical plant characteristics that can be modified include cell development (such as the number of trichomes), fruit and seed size and number, yields of plant parts such as stems, leaves and roots, the stability of the seeds during storage, characteristics of the seed pod (e.g., susceptibility to shattering), root hair length and quantity, internode distances, or the quality of seed coat. Plant growth characteristics that can be  
25 modified include growth rate, germination rate of seeds, vigor of plants and seedlings, leaf and flower senescence, male sterility, apomixis, flowering time, flower abscission, rate of nitrogen uptake, biomass or transpiration characteristics, as well as plant architecture characteristics such as apical dominance, branching patterns, number of organs, organ identity, organ shape or size.

30 **POLYPEPTIDES AND POLYNUCLEOTIDES OF THE INVENTION**

The present invention provides, among other things, transcription factors (TFs), and transcription factor homologue polypeptides, and isolated or recombinant polynucleotides encoding the polypeptides. These polypeptides and polynucleotides may be employed to modify a plant's pathogen tolerance.

35 Exemplary polynucleotides encoding the polypeptides of the invention were identified in the *Arabidopsis thaliana* GenBank database using publicly available sequence

analysis programs and parameters. Sequences initially identified were then further characterized to identify sequences comprising specified sequence strings corresponding to sequence motifs present in families of known transcription factors. Polynucleotide sequences meeting such criteria were confirmed as transcription factors.

5        Additional polynucleotides of the invention were identified by screening *Arabidopsis thaliana* and/or other plant cDNA libraries with probes corresponding to known transcription factors under low stringency hybridization conditions. Additional sequences, including full length coding sequences were subsequently recovered by the rapid amplification of cDNA ends (RACE) procedure, using a commercially available kit according  
10      to the manufacturer's instructions. Where necessary, multiple rounds of RACE are performed to isolate 5' and 3' ends. The full length cDNA was then recovered by a routine end-to-end polymerase chain reaction (PCR) using primers specific to the isolated 5' and 3' ends. Exemplary sequences are provided in the Sequence Listing.

15      The polynucleotides of the invention were ectopically expressed in overexpressor or knockout plants and changes in the pathogen tolerance of the plants was observed. Therefore, the polynucleotides and polypeptides can be employed to improve the pathogen resistance of plants.

Making polynucleotides

20      The polynucleotides of the invention include sequences that encode transcription factors and transcription factor homologue polypeptides and sequences complementary thereto, as well as unique fragments of coding sequence, or sequence complementary thereto. Such polynucleotides can be, e.g., DNA or RNA, e.g., mRNA, cRNA, synthetic RNA, genomic DNA, cDNA synthetic DNA, oligonucleotides, etc. The polynucleotides are either double-stranded or single-stranded, and include either, or both  
25      sense (i.e., coding) sequences and antisense (i.e., non-coding, complementary) sequences. The polynucleotides include the coding sequence of a transcription factor, or transcription factor homologue polypeptide, in isolation, in combination with additional coding sequences (e.g., a purification tag, a localization signal, as a fusion-protein, as a pre-protein, or the like), in combination with non-coding sequences (e.g., introns or inteins, regulatory elements such  
30      as promoters, enhancers, terminators, and the like), and/or in a vector or host environment in which the polynucleotide encoding a transcription factor or transcription factor homologue polypeptide is an endogenous or exogenous gene.

A variety of methods exist for producing the polynucleotides of the invention. Procedures for identifying and isolating DNA clones are well known to those of skill in the art, and are described in, e.g., Berger and Kimmel, Guide to Molecular Cloning Techniques, Methods in Enzymology volume 152 Academic Press, Inc., San Diego, CA ("Berger");

Sambrook et al., Molecular Cloning - A Laboratory Manual (2nd Ed.), Vol. 1-3, Cold Spring Harbor Laboratory, Cold Spring Harbor, New York, 1989 ("Sambrook") and Current Protocols in Molecular Biology, F.M. Ausubel et al., eds., Current Protocols, a joint venture between Greene Publishing Associates, Inc. and John Wiley & Sons, Inc., (supplemented through 2000) ("Ausubel").

Alternatively, polynucleotides of the invention, can be produced by a variety of in vitro amplification methods adapted to the present invention by appropriate selection of specific or degenerate primers. Examples of protocols sufficient to direct persons of skill through in vitro amplification methods, including the polymerase chain reaction (PCR) the ligase chain reaction (LCR), Qbeta-relicase amplification and other RNA polymerase mediated techniques (e.g., NASBA), e.g., for the production of the homologous nucleic acids of the invention are found in Berger, Sambrook, and Ausubel, as well as Mullis et al., (1987) PCR Protocols A Guide to Methods and Applications (Innis et al. eds) Academic Press Inc. San Diego, CA (1990) (Innis). Improved methods for cloning in vitro amplified nucleic acids are described in Wallace et al., U.S. Pat. No. 5,426,039. Improved methods for amplifying large nucleic acids by PCR are summarized in Cheng et al. (1994) Nature 369: 684-685 and the references cited therein, in which PCR amplicons of up to 40kb are generated. One of skill will appreciate that essentially any RNA can be converted into a double stranded DNA suitable for restriction digestion, PCR expansion and sequencing using reverse transcriptase and a polymerase. See, e.g., Ausubel, Sambrook and Berger, *all supra*.

Alternatively, polynucleotides and oligonucleotides of the invention can be assembled from fragments produced by solid-phase synthesis methods. Typically, fragments of up to approximately 100 bases are individually synthesized and then enzymatically or chemically ligated to produce a desired sequence, e.g., a polynucleotide encoding all or part of a transcription factor. For example, chemical synthesis using the phosphoramidite method is described, e.g., by Beaucage et al. (1981) Tetrahedron Letters 22:1859-69; and Matthes et al. (1984) EMBO J. 3:801-5. According to such methods, oligonucleotides are synthesized, purified, annealed to their complementary strand, ligated and then optionally cloned into suitable vectors. And if so desired, the polynucleotides and polypeptides of the invention can be custom ordered from any of a number of commercial suppliers.

#### HOMOLOGOUS SEQUENCES

Sequences homologous, i.e., that share significant sequence identity or similarity, to those provided in the Sequence Listing, derived from *Arabidopsis thaliana* or from other plants of choice are also an aspect of the invention. Homologous sequences can be derived from any plant including monocots and dicots and in particular agriculturally important plant species, including but not limited to, crops such as soybean, wheat, corn,

potato, cotton, rice, oilseed rape (including canola), sunflower, alfalfa, sugarcane and turf; or fruits and vegetables, such as banana, blackberry, blueberry, strawberry, and raspberry, cantaloupe, carrot, cauliflower, coffee, cucumber, eggplant, grapes, honeydew, lettuce, mango, melon, onion, papaya, peas, peppers, pineapple, spinach, squash, sweet corn, tobacco,  
5 tomato, watermelon, rosaceous fruits (such as apple, peach, pear, cherry and plum) and vegetable brassicas (such as broccoli, cabbage, cauliflower, brussel sprouts and kohlrabi). Other crops, fruits and vegetables whose phenotype can be changed include barley, rye, millet, sorghum, currant, avocado, citrus fruits such as oranges, lemons, grapefruit and tangerines, artichoke, cherries, nuts such as the walnut and peanut, endive, leek, roots, such as  
10 arrowroot, beet, cassava, turnip, radish, yam, and sweet potato, and beans. The homologous sequences may also be derived from woody species, such pine, poplar and eucalyptus.

Transcription factors that are homologous to the listed sequences will typically share at least about 31% amino acid sequence identity. More closely related transcription factors can share at least about 50%, about 60%, about 65%, about 70%, about  
15 75% or about 80% or about 90% or about 95% or about 98% or more sequence identity with the listed sequences. Factors that are most closely related to the listed sequences share, e.g., at least about 85%, about 90% or about 95% or more % sequence identity to the listed sequences. At the nucleotide level, the sequences will typically share at least about 40% nucleotide sequence identity, preferably at least about 50%, about 60%, about 70% or about  
20 80% sequence identity, and more preferably about 85%, about 90%, about 95% or about 97% or more sequence identity to one or more of the listed sequences. The degeneracy of the genetic code enables major variations in the nucleotide sequence of a polynucleotide while maintaining the amino acid sequence of the encoded protein. Conserved domains within a transcription factor family may exhibit a higher degree of sequence homology, such as at least  
25 65% sequence identity including conservative substitutions, and preferably at least 80% sequence identity.

#### Identifying Nucleic Acids by Hybridization

Polynucleotides homologous to the sequences illustrated in the Sequence Listing can be identified, e.g., by hybridization to each other under stringent or under highly  
30 stringent conditions. Single stranded polynucleotides hybridize when they associate based on a variety of well characterized physico-chemical forces, such as hydrogen bonding, solvent exclusion, base stacking and the like. The stringency of a hybridization reflects the degree of sequence identity of the nucleic acids involved, such that the higher the stringency, the more similar are the two polynucleotide strands. Stringency is influenced by a variety of factors, including temperature, salt concentration and composition, organic and non-organic additives, solvents, etc. present in both the hybridization and wash solutions and incubations (and number), as described in more detail in the references cited above.

An example of stringent hybridization conditions for hybridization of complementary nucleic acids which have more than 100 complementary residues on a filter in a Southern or northern blot is about 5°C to 20°C lower than the thermal melting point (T<sub>m</sub>) for the specific sequence at a defined ionic strength and pH. The T<sub>m</sub> is the temperature (under defined ionic strength and pH) at which 50% of the target sequence hybridizes to a perfectly matched probe. Nucleic acid molecules that hybridize under stringent conditions will typically hybridize to a probe based on either the entire cDNA or selected portions, e.g., to a unique subsequence, of the cDNA under wash conditions of 0.2x SSC to 2.0 x SSC, 0.1% SDS at 50-65° C, for example 0.2 x SSC, 0.1% SDS at 65° C. For identification of less closely related homologues washes can be performed at a lower temperature, e.g., 50° C. In general, stringency is increased by raising the wash temperature and/or decreasing the concentration of SSC.

As another example, stringent conditions can be selected such that an oligonucleotide that is perfectly complementary to the coding oligonucleotide hybridizes to the coding oligonucleotide with at least about a 5-10x higher signal to noise ratio than the ratio for hybridization of the perfectly complementary oligonucleotide to a nucleic acid encoding a transcription factor known as of the filing date of the application. Conditions can be selected such that a higher signal to noise ratio is observed in the particular assay which is used, e.g., about 15x, 25x, 35x, 50x or more. Accordingly, the subject nucleic acid hybridizes to the unique coding oligonucleotide with at least a 2x higher signal to noise ratio as compared to hybridization of the coding oligonucleotide to a nucleic acid encoding known polypeptide. Again, higher signal to noise ratios can be selected, e.g., about 5x, 10x, 25x, 35x, 50x or more. The particular signal will depend on the label used in the relevant assay, e.g., a fluorescent label, a colorimetric label, a radio active label, or the like.

Alternatively, transcription factor homologue polypeptides can be obtained by screening an expression library using antibodies specific for one or more transcription factors. With the provision herein of the disclosed transcription factor, and transcription factor homologue nucleic acid sequences, the encoded polypeptide(s) can be expressed and purified in a heterologous expression system (e.g., *E. coli*) and used to raise antibodies (monoclonal or polyclonal) specific for the polypeptide(s) in question. Antibodies can also be raised against synthetic peptides derived from transcription factor, or transcription factor homologue, amino acid sequences. Methods of raising antibodies are well known in the art and are described in Harlow and Lane (1988) Antibodies: A Laboratory Manual, Cold Spring Harbor Laboratory, New York. Such antibodies can then be used to screen an expression library produced from the plant from which it is desired to clone additional transcription

factor homologues, using the methods described above. The selected cDNAs can be confirmed by sequencing and enzymatic activity.

#### SEQUENCE VARIATIONS

It will readily be appreciated by those of skill in the art, that any of a variety 5 of polynucleotide sequences are capable of encoding the transcription factors and transcription factor homologue polypeptides of the invention. Due to the degeneracy of the genetic code, many different polynucleotides can encode identical and/or substantially similar polypeptides in addition to those sequences illustrated in the Sequence Listing.

For example, Table 1 illustrates, e.g., that the codons AGC, AGT, TCA, 10 TCC, TCG, and TCT all encode the same amino acid: serine. Accordingly, at each position in the sequence where there is a codon encoding serine, any of the above trinucleotide sequences can be used without altering the encoded polypeptide.

Table 1

Amino acids			Codon				
Alanine	Ala	A	GCA	GCC	GCG	GCU	
Cysteine	Cys	C	TGC	TGT			
Aspartic acid	Asp	D	GAC	GAT			
Glutamic acid	Glu	E	GAA	GAG			
Phenylalanine	Phe	F	TTC	TTT			
Glycine	Gly	G	GGA	GGC	GGG	GGT	
Histidine	His	H	CAC	CAT			
Isoleucine	Ile	I	ATA	ATC	ATT		
Lysine	Lys	K	AAA	AAG			
Leucine	Leu	L	TTA	TTG	CTA	CTC	CTG CTT
Methionine	Met	M	ATG				
Asparagine	Asn	N	AAC	AAT			
Proline	Pro	P	CCA	CCC	CCG	CCT	
Glutamine	Gln	Q	CAA	CAG			
Arginine	Arg	R	AGA	AGG	CGA	CGC	CGG CGT
Serine	Ser	S	AGC	AGT	TCA	TCC	TCG TCT
Threonine	Thr	T	ACA	ACC	ACG	ACT	
Valine	Val	V	GTA	GTC	GTG	GTT	
Tryptophan	Trp	W	TGG				
Tyrosine	Tyr	Y	TAC	TAT			

15

Sequence alterations that do not change the amino acid sequence encoded by the polynucleotide are termed "silent" variations. With the exception of the codons ATG and TGG, encoding methionine and tryptophan, respectively, any of the possible codons for the same amino acid can be substituted by a variety of techniques, e.g., site-directed mutagenesis, 20 available in the art. Accordingly, any and all such variations of a sequence selected from the above table are a feature of the invention.

In addition to silent variations, other conservative variations that alter one, or a few amino acids in the encoded polypeptide, can be made without altering the function of the polypeptide, these conservative variants are, likewise, a feature of the invention.

For example, substitutions, deletions and insertions introduced into the  
5 sequences provided in the Sequence Listing are also envisioned by the invention. Such sequence modifications can be engineered into a sequence by site-directed mutagenesis (Wu (ed.) Meth. Enzymol. (1993) vol. 217, Academic Press) or the other methods noted below. Amino acid substitutions are typically of single residues; insertions usually will be on the  
10 order of about from 1 to 10 amino acid residues; and deletions will range about from 1 to 30 residues. In preferred embodiments, deletions or insertions are made in adjacent pairs, e.g., a deletion of two residues or insertion of two residues. Substitutions, deletions, insertions or any combination thereof can be combined to arrive at a sequence. The mutations that are made in the polynucleotide encoding the transcription factor should not place the sequence out of reading frame and should not create complementary regions that could produce  
15 secondary mRNA structure. Preferably, the polypeptide encoded by the DNA performs the desired function.

Conservative substitutions are those in which at least one residue in the amino acid sequence has been removed and a different residue inserted in its place. Such substitutions generally are made in accordance with the Table 2 when it is desired to maintain  
20 the activity of the protein. Table 2 shows amino acids which can be substituted for an amino acid in a protein and which are typically regarded as conservative substitutions.

25

30

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Table 2

Residue	Conservative Substitutions
Ala	Ser
Arg	Lys
Asn	Gln; His
Asp	Glu
Gln	Asn
Cys	Ser
Glu	Asp
Gly	Pro
His	Asn; Gln
Ile	Leu, Val
Leu	Ile; Val
Lys	Arg; Gln
Met	Leu; Ile
Phe	Met; Leu; Tyr
Ser	Thr; Gly
Thr	Ser; Val
Trp	Tyr
Tyr	Trp; Phe
Val	Ile; Leu

Substitutions that are less conservative than those in Table 2 can be selected by picking residues that differ more significantly in their effect on maintaining (a) the structure of the polypeptide backbone in the area of the substitution, for example, as a sheet or helical conformation, (b) the charge or hydrophobicity of the molecule at the target site, or (c) the bulk of the side chain. The substitutions which in general are expected to produce the greatest changes in protein properties will be those in which (a) a hydrophilic residue, e.g., seryl or threonyl, is substituted for (or by) a hydrophobic residue, e.g., leucyl, isoleucyl, phenylalanyl, valyl or alanyl; (b) a cysteine or proline is substituted for (or by) any other residue; (c) a residue having an electropositive side chain, e.g., lysyl, arginyl, or histidyl, is substituted for (or by) an electronegative residue, e.g., glutamyl or aspartyl; or (d) a residue having a bulky side chain, e.g., phenylalanine, is substituted for (or by) one not having a side chain, e.g., glycine.

FURTHER MODIFYING SEQUENCES OF THE INVENTION—MUTATION/ FORCED EVOLUTION

5 In addition to generating silent or conservative substitutions as noted, above, the present invention optionally includes methods of modifying the sequences of the Sequence Listing. In the methods, nucleic acid or protein modification methods are used to alter the given sequences to produce new sequences and/or to chemically or enzymatically modify given sequences to change the properties of the nucleic acids or proteins.

10 Thus, in one embodiment, given nucleic acid sequences are modified, e.g., according to standard mutagenesis or artificial evolution methods to produce modified sequences. For example, Ausubel, *supra*, provides additional details on mutagenesis methods. Artificial forced evolution methods are described, e.g., by Stemmer (1994) *Nature* 370:389-391, and Stemmer (1994) *Proc. Natl. Acad. Sci. USA* 91:10747-10751. Many other mutation and evolution methods are also available and expected to be within the skill of the practitioner.

15 Similarly, chemical or enzymatic alteration of expressed nucleic acids and polypeptides can be performed by standard methods. For example, sequence can be modified by addition of lipids, sugars, peptides, organic or inorganic compounds, by the inclusion of modified nucleotides or amino acids, or the like. For example, protein modification techniques are illustrated in Ausubel, *supra*. Further details on chemical and enzymatic modifications can be found herein. These modification methods can be used to modify any given sequence, or to modify any sequence produced by the various mutation and artificial evolution modification methods noted herein.

20 Accordingly, the invention provides for modification of any given nucleic acid by mutation, evolution, chemical or enzymatic modification, or other available methods, as well as for the products produced by practicing such methods, e.g., using the sequences herein as a starting substrate for the various modification approaches.

25 For example, optimized coding sequence containing codons preferred by a particular prokaryotic or eukaryotic host can be used e.g., to increase the rate of translation or to produce recombinant RNA transcripts having desirable properties, such as a longer half-life, as compared with transcripts produced using a non-optimized sequence. Translation stop codons can also be modified to reflect host preference. For example, preferred stop codons for *S. cerevisiae* and mammals are TAA and TGA, respectively. The preferred stop codon for monocotyledonous plants is TGA, whereas insects and *E. coli* prefer to use TAA as the stop codon.

30 The polynucleotide sequences of the present invention can also be engineered in order to alter a coding sequence for a variety of reasons, including but not limited to, alterations which modify the sequence to facilitate cloning, processing and/or expression of

the gene product. For example, alterations are optionally introduced using techniques which are well known in the art, e.g., site-directed mutagenesis, to insert new restriction sites, to alter glycosylation patterns, to change codon preference, to introduce splice sites, etc.

Furthermore, a fragment or domain derived from any of the polypeptides of  
5 the invention can be combined with domains derived from other transcription factors or synthetic domains to modify the biological activity of a transcription factor. For instance, a DNA binding domain derived from a transcription factor of the invention can be combined with the activation domain of another transcription factor or with a synthetic activation domain. A transcription activation domain assists in initiating transcription from a DNA  
10 binding site. Examples include the transcription activation region of VP16 or GAL4 (Moore et al. (1998) Proc. Natl. Acad. Sci. USA 95: 376-381; and Aoyama et al. (1995) Plant Cell 7:1773-1785), peptides derived from bacterial sequences (Ma and Ptashne (1987) Cell 51; 113-119) and synthetic peptides (Giniger and Ptashne, (1987) Nature 330:670-672).

#### EXPRESSION AND MODIFICATION OF POLYPEPTIDES

15 Typically, polynucleotide sequences of the invention are incorporated into recombinant DNA (or RNA) molecules that direct expression of polypeptides of the invention in appropriate host cells, transgenic plants, in vitro translation systems, or the like. Due to the inherent degeneracy of the genetic code, nucleic acid sequences which encode substantially the same or a functionally equivalent amino acid sequence can be substituted for any listed  
20 sequence to provide for cloning and expressing the relevant homologue.

##### Vectors, Promoters and Expression Systems

The present invention includes recombinant constructs comprising one or more of the nucleic acid sequences herein. The constructs typically comprise a vector, such as a plasmid, a cosmid, a phage, a virus (e.g., a plant virus), a bacterial artificial chromosome (BAC), a yeast artificial chromosome (YAC), or the like, into which a nucleic acid sequence of the invention has been inserted, in a forward or reverse orientation. In a preferred aspect of this embodiment, the construct further comprises regulatory sequences, including, for example, a promoter, operably linked to the sequence. Large numbers of suitable vectors and promoters are known to those of skill in the art, and are commercially available.  
25

30 General texts which describe molecular biological techniques useful herein, including the use and production of vectors, promoters and many other relevant topics, include Berger, Sambrook and Ausubel, *supra*. Any of the identified sequences can be incorporated into a cassette or vector, e.g., for expression in plants. A number of expression vectors suitable for stable transformation of plant cells or for the establishment of transgenic  
35 plants have been described including those described in Weissbach and Weissbach, (1989) Methods for Plant Molecular Biology, Academic Press, and Gelvin et al., (1990) Plant

Molecular Biology Manual, Kluwer Academic Publishers. Specific examples include those derived from a Ti plasmid of *Agrobacterium tumefaciens*, as well as those disclosed by Herrera-Estrella et al. (1983) Nature 303: 209, Bevan (1984) Nucl Acid Res. 12: 8711-8721, Klee (1985) Bio/Technology 3: 637-642, for dicotyledonous plants.

5        Alternatively, non-Ti vectors can be used to transfer the DNA into monocotyledonous plants and cells by using free DNA delivery techniques. Such methods can involve, for example, the use of liposomes, electroporation, microprojectile bombardment, silicon carbide whiskers, and viruses. By using these methods transgenic plants such as wheat, rice (Christou (1991) Bio/Technology 9: 957-962) and corn (Gordon-  
10 Kamm (1990) Plant Cell 2: 603-618) can be produced. An immature embryo can also be a good target tissue for monocots for direct DNA delivery techniques by using the particle gun (Weeks et al. (1993) Plant Physiol 102: 1077-1084; Vasil (1993) Bio/Technology 10: 667-  
15 674; Wan and Lemeaux (1994) Plant Physiol 104: 37-48, and for *Agrobacterium*-mediated DNA transfer (Ishida et al. (1996) Nature Biotech 14: 745-750).

15        Typically, plant transformation vectors include one or more cloned plant coding sequence (genomic or cDNA) under the transcriptional control of 5' and 3' regulatory sequences and a dominant selectable marker. Such plant transformation vectors typically also contain a promoter (e.g., a regulatory region controlling inducible or constitutive, environmentally-or developmentally-regulated, or cell- or tissue-specific expression), a  
20 transcription initiation start site, an RNA processing signal (such as intron splice sites), a transcription termination site, and/or a polyadenylation signal.

25        Examples of constitutive plant promoters which can be useful for expressing the TF sequence include: the cauliflower mosaic virus (CaMV) 35S promoter, which confers constitutive, high-level expression in most plant tissues (see, e.g., Odel et al. (1985) Nature 313:810); the nopaline synthase promoter (An et al. (1988) Plant Physiol 88:547); and the octopine synthase promoter (Fromm et al. (1989) Plant Cell 1: 977).

30        A variety of plant gene promoters that regulate gene expression in response to environmental, hormonal, chemical, developmental signals, and in a tissue-active manner can be used for expression of a TF sequence in plants. Choice of a promoter is based largely on the phenotype of interest and is determined by such factors as tissue (e.g., seed, fruit, root, pollen, vascular tissue, flower, carpel, etc.), inducibility (e.g., in response to wounding, heat, cold, drought, light, pathogens, etc.), timing, developmental stage, and the like. Numerous known promoters have been characterized and can favorable be employed to promote expression of a polynucleotide of the invention in a transgenic plant or cell of interest. For  
35 example, tissue specific promoters include: seed-specific promoters (such as the napin, phaseolin or DC3 promoter described in US Pat. No. 5,773,697), fruit-specific promoters that are active during fruit ripening (such as the dru 1 promoter (US Pat. No. 5,783,393), or the

2A11 promoter (US Pat. No. 4,943,674) and the tomato polygalacturonase promoter (Bird et al. (1988) *Plant Mol Biol* 11:651), root-specific promoters, such as those disclosed in US Patent Nos. 5,618,988, 5,837,848 and 5,905,186, pollen-active promoters such as PTA29, PTA26 and PTA13 (US Pat. No. 5,792,929), promoters active in vascular tissue (Ringli and Keller (1998) *Plant Mol Biol* 37:977-988), flower-specific (Kaiser et al, (1995) *Plant Mol Biol* 28:231-243), pollen (Baerson et al. (1994) *Plant Mol Biol* 26:1947-1959), carpels (Ohl et al. (1990) *Plant Cell* 2:837-848), pollen and ovules (Baerson et al. (1993) *Plant Mol Biol* 22:255-267), auxin-inducible promoters (such as that described in van der Kop et al. (1999) *Plant Mol Biol* 39:979-990 or Baumann et al. (1999) *Plant Cell* 11:323-334), cytokinin-inducible promoter (Guevara-Garcia (1998) *Plant Mol Biol* 38:743-753), promoters responsive to gibberellin (Shi et al. (1998) *Plant Mol Biol* 38:1053-1060, Willmott et al. (1998) 38:817-825) and the like. Additional promoters are those that elicit expression in response to heat (Ainley et al. (1993) *Plant Mol Biol* 22: 13-23), light (e.g., the pea rbcS-3A promoter, Kuhlemeier et al. (1989) *Plant Cell* 1:471, and the maize rbcS promoter, Schaffner and Sheen (1991) *Plant Cell* 3: 997); wounding (e.g., *wunI*, Siebertz et al. (1989) *Plant Cell* 1: 961); pathogens (such as the PR-1 promoter described in Buchel et al. (1999) *Plant Mol. Biol.* 40:387-396, and the PDF1.2 promoter described in Manners et al. (1998) *Plant Mol. Biol.* 38:1071-80), and chemicals such as methyl jasmonate or salicylic acid (Gatz et al. (1997) *Plant Mol Biol* 48: 89-108). In addition, the timing of the expression can be controlled by using promoters such as those acting at senescence (An and Amazon (1995) *Science* 270: 1986-1988); or late seed development (Odell et al. (1994) *Plant Physiol* 106:447-458).

Plant expression vectors can also include RNA processing signals that can be positioned within, upstream or downstream of the coding sequence. In addition, the expression vectors can include additional regulatory sequences from the 3'-untranslated region of plant genes, e.g., a 3' terminator region to increase mRNA stability of the mRNA, such as the PI-II terminator region of potato or the octopine or nopaline synthase 3' terminator regions.

#### Additional Expression Elements

Specific initiation signals can aid in efficient translation of coding sequences. These signals can include, e.g., the ATG initiation codon and adjacent sequences. In cases where a coding sequence, its initiation codon and upstream sequences are inserted into the appropriate expression vector, no additional translational control signals may be needed. However, in cases where only coding sequence (e.g., a mature protein coding sequence), or a portion thereof, is inserted, exogenous transcriptional control signals including the ATG initiation codon can be separately provided. The initiation codon is provided in the correct reading frame to facilitate transcription. Exogenous transcriptional elements and initiation

codons can be of various origins, both natural and synthetic. The efficiency of expression can be enhanced by the inclusion of enhancers appropriate to the cell system in use.

Expression Hosts

The present invention also relates to host cells which are transduced with vectors of the invention, and the production of polypeptides of the invention (including fragments thereof) by recombinant techniques. Host cells are genetically engineered (i.e., nucleic acids are introduced, e.g., transduced, transformed or transfected) with the vectors of this invention, which may be, for example, a cloning vector or an expression vector comprising the relevant nucleic acids herein. The vector is optionally a plasmid, a viral particle, a phage, a naked nucleic acids, etc. The engineered host cells can be cultured in conventional nutrient media modified as appropriate for activating promoters, selecting transformants, or amplifying the relevant gene. The culture conditions, such as temperature, pH and the like, are those previously used with the host cell selected for expression, and will be apparent to those skilled in the art and in the references cited herein, including, Sambrook and Ausubel.

The host cell can be a eukaryotic cell, such as a yeast cell, or a plant cell, or the host cell can be a prokaryotic cell, such as a bacterial cell. Plant protoplasts are also suitable for some applications. For example, the DNA fragments are introduced into plant tissues, cultured plant cells or plant protoplasts by standard methods including electroporation (Fromm et al., (1985) Proc. Natl. Acad. Sci. USA 82, 5824, infection by viral vectors such as cauliflower mosaic virus (CaMV) (Hohn et al., (1982) Molecular Biology of Plant Tumors, (Academic Press, New York) pp. 549-560; US 4,407,956), high velocity ballistic penetration by small particles with the nucleic acid either within the matrix of small beads or particles, or on the surface (Klein et al., (1987) Nature 327, 70-73), use of pollen as vector (WO 85/01856), or use of *Agrobacterium tumefaciens* or *A. rhizogenes* carrying a T-DNA plasmid in which DNA fragments are cloned. The T-DNA plasmid is transmitted to plant cells upon infection by *Agrobacterium tumefaciens*, and a portion is stably integrated into the plant genome (Horsch et al. (1984) Science 233:496-498; Fraley et al. (1983) Proc. Natl. Acad. Sci. USA 80, 4803).

The cell can include a nucleic acid of the invention which encodes a polypeptide, wherein the cells expresses a polypeptide of the invention. The cell can also include vector sequences, or the like. Furthermore, cells and transgenic plants which include any polypeptide or nucleic acid above or throughout this specification, e.g., produced by transduction of a vector of the invention, are an additional feature of the invention.

For long-term, high-yield production of recombinant proteins, stable expression can be used. Host cells transformed with a nucleotide sequence encoding a polypeptide of the invention are optionally cultured under conditions suitable for the

expression and recovery of the encoded protein from cell culture. The protein or fragment thereof produced by a recombinant cell may be secreted, membrane-bound, or contained intracellularly, depending on the sequence and/or the vector used. As will be understood by those of skill in the art, expression vectors containing polynucleotides encoding mature proteins of the invention can be designed with signal sequences which direct secretion of the mature polypeptides through a prokaryotic or eukaryotic cell membrane.

**Modified Amino Acids**

10 Polypeptides of the invention may contain one or more modified amino acids. The presence of modified amino acids may be advantageous in, for example, increasing polypeptide half-life, reducing polypeptide antigenicity or toxicity, increasing polypeptide storage stability, or the like. Amino acid(s) are modified, for example, co-translationally or post-translationally during recombinant production or modified by synthetic or chemical means.

15 Non-limiting examples of a modified amino acid include incorporation or other use of acetylated amino acids, glycosylated amino acids, sulfated amino acids, prenylated (e.g., farnesylated, geranylgeranylated) amino acids, PEG modified (e.g., "PEGylated") amino acids, biotinylated amino acids, carboxylated amino acids, phosphorylated amino acids, etc. References adequate to guide one of skill in the modification of amino acids are replete throughout the literature.

20 **IDENTIFICATION OF ADDITIONAL FACTORS**

A transcription factor provided by the present invention can also be used to identify additional endogenous or exogenous molecules that can affect a phenotype or trait of interest. On the one hand, such molecules include organic (small or large molecules) and/or inorganic compounds that affect expression of (i.e., regulate) a particular transcription factor. 25 Alternatively, such molecules include endogenous molecules that are acted upon either at a transcriptional level by a transcription factor of the invention to modify a phenotype as desired. For example, the transcription factors can be employed to identify one or more downstream gene with which is subject to a regulatory effect of the transcription factor. In one approach, a transcription factor or transcription factor homologue of the invention is expressed in a host cell, e.g., a transgenic plant cell, tissue or explant, and expression products, either RNA or protein, of likely or random targets are monitored, e.g., by hybridization to a microarray of nucleic acid probes corresponding to genes expressed in a tissue or cell type of interest, by two-dimensional gel electrophoresis of protein products, or by any other method known in the art for assessing expression of gene products at the level of RNA or protein. 30 35 Alternatively, a transcription factor of the invention can be used to identify promoter sequences (i.e., binding sites) involved in the regulation of a downstream target. After

identifying a promoter sequence, interactions between the transcription factor and the promoter sequence can be modified by changing specific nucleotides in the promoter sequence or specific amino acids in the transcription factor that interact with the promoter sequence to alter a plant trait. Typically, transcription factor DNA binding sites are identified  
5 by gel shift assays. After identifying the promoter regions, the promoter region sequences can be employed in double-stranded DNA arrays to identify molecules that affect the interactions of the transcription factors with their promoters (Bulyk et al. (1999) *Nature Biotechnology* 17:573-577).

The identified transcription factors are also useful to identify proteins that  
10 modify the activity of the transcription factor. Such modification can occur by covalent modification, such as by phosphorylation, or by protein-protein (homo or-heteropolymer) interactions. Any method suitable for detecting protein-protein interactions can be employed. Among the methods that can be employed are co-immunoprecipitation, cross-linking and copurification through gradients or chromatographic columns, and the two-hybrid yeast system.  
15

The two-hybrid system detects protein interactions *in vivo* and is described in Chien, et al., (1991), *Proc. Natl. Acad. Sci. USA* 88, 9578-9582 and is commercially available from Clontech (Palo Alto, Calif.). In such a system, plasmids are constructed that encode two hybrid proteins: one consists of the DNA-binding domain of a transcription activator protein fused to the TF polypeptide and the other consists of the transcription activator protein's activation domain fused to an unknown protein that is encoded by a cDNA that has been recombined into the plasmid as part of a cDNA library. The DNA-binding domain fusion plasmid and the cDNA library are transformed into a strain of the yeast *Saccharomyces cerevisiae* that contains a reporter gene (e.g., lacZ) whose regulatory region contains the transcription activator's binding site. Either hybrid protein alone cannot activate transcription  
20 of the reporter gene. Interaction of the two hybrid proteins reconstitutes the functional activator protein and results in expression of the reporter gene, which is detected by an assay for the reporter gene product. Then, the library plasmids responsible for reporter gene expression are isolated and sequenced to identify the proteins encoded by the library plasmids. After identifying proteins that interact with the transcription factors, assays for  
25 compounds that interfere with the TF protein-protein interactions can be performed.  
30

#### IDENTIFICATION OF MODULATORS

In addition to the intracellular molecules described above, extracellular molecules that alter activity or expression of a transcription factor, either directly or indirectly, can be identified. For example, the methods can entail first placing a candidate  
35 molecule in contact with a plant or plant cell. The molecule can be introduced by topical administration, such as spraying or soaking of a plant, and then the molecule's effect on the

expression or activity of the TF polypeptide or the expression of the polynucleotide monitored. Changes in the expression of the TF polypeptide can be monitored by use of polyclonal or monoclonal antibodies, gel electrophoresis or the like. Changes in the expression of the corresponding polynucleotide sequence can be detected by use of microarrays, Northerns, quantitative PCR, or any other technique for monitoring changes in mRNA expression. These techniques are exemplified in Ausubel et al. (eds) Current Protocols in Molecular Biology, John Wiley & Sons (1998). Such changes in the expression levels can be correlated with modified plant traits and thus identified molecules can be useful for soaking or spraying on fruit, vegetable and grain crops to modify traits in plants.

Essentially any available composition can be tested for modulatory activity of expression or activity of any nucleic acid or polypeptide herein. Thus, available libraries of compounds such as chemicals, polypeptides, nucleic acids and the like can be tested for modulatory activity. Often, potential modulator compounds can be dissolved in aqueous or organic (e.g., DMSO-based) solutions for easy delivery to the cell or plant of interest in which the activity of the modulator is to be tested. Optionally, the assays are designed to screen large modulator composition libraries by automating the assay steps and providing compounds from any convenient source to assays, which are typically run in parallel (e.g., in microtiter formats on microtiter plates in robotic assays).

In one embodiment, high throughput screening methods involve providing a combinatorial library containing a large number of potential compounds (potential modulator compounds). Such "combinatorial chemical libraries" are then screened in one or more assays, as described herein, to identify those library members (particular chemical species or subclasses) that display a desired characteristic activity. The compounds thus identified can serve as target compounds.

A combinatorial chemical library can be, e.g., a collection of diverse chemical compounds generated by chemical synthesis or biological synthesis. For example, a combinatorial chemical library such as a polypeptide library is formed by combining a set of chemical building blocks (e.g., in one example, amino acids) in every possible way for a given compound length (i.e., the number of amino acids in a polypeptide compound of a set length). Exemplary libraries include peptide libraries, nucleic acid libraries, antibody libraries (see, e.g., Vaughn et al. (1996) Nature Biotechnology, 14(3):309-314 and PCT/US96/10287), carbohydrate libraries (see, e.g., Liang et al. Science (1996) 274:1520-1522 and U.S. Patent 5,593,853), peptide nucleic acid libraries (see, e.g., U.S. Patent 5,539,083), and small organic molecule libraries (see, e.g., benzodiazepines, Baum C&EN Jan 18, page 33 (1993); isoprenoids, U.S. Patent 5,569,588; thiazolidinones and metathiazanones, U.S. Patent 5,549,974; pyrrolidines, U.S. Patents 5,525,735 and 5,519,134; morpholino compounds, U.S. Patent 5,506,337) and the like.

Preparation and screening of combinatorial or other libraries is well known to those of skill in the art. Such combinatorial chemical libraries include, but are not limited to, peptide libraries (see, e.g., U.S. Patent 5,010,175, Furka, *Int. J. Pept. Prot. Res.* 37:487-493 (1991) and Houghton et al. *Nature* 354:84-88 (1991)). Other chemistries for generating 5 chemical diversity libraries can also be used.

In addition, as noted, compound screening equipment for high-throughput screening is generally available, e.g., using any of a number of well known robotic systems that have also been developed for solution phase chemistries useful in assay systems. These 10 systems include automated workstations including an automated synthesis apparatus and robotic systems utilizing robotic arms. Any of the above devices are suitable for use with the present invention, e.g., for high-throughput screening of potential modulators. The nature and implementation of modifications to these devices (if any) so that they can operate as discussed herein will be apparent to persons skilled in the relevant art.

Indeed, entire high throughput screening systems are commercially available. 15 These systems typically automate entire procedures including all sample and reagent pipetting, liquid dispensing, timed incubations, and final readings of the microplate in detector(s) appropriate for the assay. These configurable systems provide high throughput and rapid start up as well as a high degree of flexibility and customization. Similarly, microfluidic implementations of screening are also commercially available.

The manufacturers of such systems provide detailed protocols the various 20 high throughput. Thus, for example, Zymark Corp. provides technical bulletins describing screening systems for detecting the modulation of gene transcription, ligand binding, and the like. The integrated systems herein, in addition to providing for sequence alignment and, optionally, synthesis of relevant nucleic acids, can include such screening apparatus to 25 identify modulators that have an effect on one or more polynucleotides or polypeptides according to the present invention.

In some assays it is desirable to have positive controls to ensure that the components of the assays are working properly. At least two types of positive controls are appropriate. That is, known transcriptional activators or inhibitors can be incubated with 30 cells/plants/ etc. in one sample of the assay, and the resulting increase/decrease in transcription can be detected by measuring the resulting increase in RNA/ protein expression, etc., according to the methods herein. It will be appreciated that modulators can also be combined with transcriptional activators or inhibitors to find modulators which inhibit transcriptional activation or transcriptional repression. Either expression of the nucleic acids 35 and proteins herein or any additional nucleic acids or proteins activated by the nucleic acids or proteins herein, or both, can be monitored.

In an embodiment, the invention provides a method for identifying compositions that modulate the activity or expression of a polynucleotide or polypeptide of the invention. For example, a test compound, whether a small or large molecule, is placed in contact with a cell, plant (or plant tissue or explant), or composition comprising the 5 polynucleotide or polypeptide of interest and a resulting effect on the cell, plant, (or tissue or explant) or composition is evaluated by monitoring, either directly or indirectly, one or more of: expression level of the polynucleotide or polypeptide, activity (or modulation of the activity) of the polynucleotide or polypeptide. In some cases, an alteration in a plant phenotype can be detected following contact of a plant (or plant cell, or tissue or explant) with 10 the putative modulator, e.g., by modulation of expression or activity of a polynucleotide or polypeptide of the invention.

#### SUBSEQUENCES

Also contemplated are uses of polynucleotides, also referred to herein as 15 oligonucleotides, typically having at least 12 bases, preferably at least 15, more preferably at least 20, 30, or 50 bases, which hybridize under at least highly stringent (or ultra-high stringent or ultra-ultra- high stringent conditions) conditions to a polynucleotide sequence described above. The polynucleotides may be used as probes, primers, sense and antisense agents, and the like, according to methods as noted *supra*.

Subsequences of the polynucleotides of the invention, including 20 polynucleotide fragments and oligonucleotides are useful as nucleic acid probes and primers. An oligonucleotide suitable for use as a probe or primer is at least about 15 nucleotides in length, more often at least about 18 nucleotides, often at least about 21 nucleotides, frequently at least about 30 nucleotides, or about 40 nucleotides, or more in length. A nucleic acid probe 25 is useful in hybridization protocols, e.g., to identify additional polypeptide homologues of the invention, including protocols for microarray experiments. Primers can be annealed to a complementary target DNA strand by nucleic acid hybridization to form a hybrid between the primer and the target DNA strand, and then extended along the target DNA strand by a DNA polymerase enzyme. Primer pairs can be used for amplification of a nucleic acid sequence, 30 e.g., by the polymerase chain reaction (PCR) or other nucleic-acid amplification methods. See Sambrook and Ausubel, *supra*.

In addition, the invention includes an isolated or recombinant polypeptide 35 including a subsequence of at least about 15 contiguous amino acids encoded by the recombinant or isolated polynucleotides of the invention. For example, such polypeptides, or domains or fragments thereof, can be used as immunogens, e.g., to produce antibodies specific for the polypeptide sequence, or as probes for detecting a sequence of interest. A

subsequence can range in size from about 15 amino acids in length up to and including the full length of the polypeptide.

### PRODUCTION OF TRANSGENIC PLANTS

#### Modification of Traits

5       The polynucleotides of the invention are favorably employed to produce transgenic plants with various traits, or characteristics, that have been modified in a desirable manner, e.g., to improve the pathogen resistance of a plant. For example, alteration of expression levels or patterns (e.g., spatial or temporal expression patterns) of one or more of the transcription factors (or transcription factor homologues) of the invention, as compared with the levels of the same protein found in a wild type plant, can be used to modify a plant's traits. An illustrative example of trait modification, improved pathogen tolerance, by altering expression levels of a particular transcription factor is described further in the Examples and the Sequence Listing.

#### Antisense and Cosuppression Approaches

15      In addition to expression of the nucleic acids of the invention as gene replacement or plant phenotype modification nucleic acids, the nucleic acids are also useful for sense and anti-sense suppression of expression, e.g., to down-regulate expression of a nucleic acid of the invention, e.g., as a further mechanism for modulating plant phenotype. That is, the nucleic acids of the invention, or subsequences or anti-sense sequences thereof, 20 can be used to block expression of naturally occurring homologous nucleic acids. A variety of sense and anti-sense technologies are known in the art, e.g., as set forth in Lichtenstein and Nellen (1997) Antisense Technology: A Practical Approach IRL Press at Oxford University, Oxford, England. In general, sense or anti-sense sequences are introduced into a cell, where they are optionally amplified, e.g., by transcription. Such sequences include both simple 25 oligonucleotide sequences and catalytic sequences such as ribozymes.

For example, a reduction or elimination of expression (i.e., a "knock-out") of a transcription factor or transcription factor homologue polypeptide in a transgenic plant, e.g., to modify a plant trait, can be obtained by introducing an antisense construct corresponding to the polypeptide of interest as a cDNA. For antisense suppression, the transcription factor or 30 homologue cDNA is arranged in reverse orientation (with respect to the coding sequence) relative to the promoter sequence in the expression vector. The introduced sequence need not be the full length cDNA or gene, and need not be identical to the cDNA or gene found in the plant type to be transformed. Typically, the antisense sequence need only be capable of hybridizing to the target gene or RNA of interest. Thus, where the introduced sequence is of 35 shorter length, a higher degree of homology to the endogenous transcription factor sequence will be needed for effective antisense suppression. While antisense sequences of various

lengths can be utilized, preferably, the introduced antisense sequence in the vector will be at least 30 nucleotides in length, and improved antisense suppression will typically be observed as the length of the antisense sequence increases. Preferably, the length of the antisense sequence in the vector will be greater than 100 nucleotides. Transcription of an antisense construct as described results in the production of RNA molecules that are the reverse complement of mRNA molecules transcribed from the endogenous transcription factor gene in the plant cell.

Suppression of endogenous transcription factor gene expression can also be achieved using a ribozyme. Ribozymes are RNA molecules that possess highly specific 10 endoribonuclease activity. The production and use of ribozymes are disclosed in U.S. Patent No. 4,987,071 and U.S. Patent No. 5,543,508. Synthetic ribozyme sequences including antisense RNAs can be used to confer RNA cleaving activity on the antisense RNA, such that endogenous mRNA molecules that hybridize to the antisense RNA are cleaved, which in turn leads to an enhanced antisense inhibition of endogenous gene expression.

Vectors in which RNA encoded by a transcription factor or transcription factor homologue cDNA is over-expressed can also be used to obtain co-suppression of a corresponding endogenous gene, e.g., in the manner described in U.S. Patent No. 5,231,020 to Jorgensen. Such co-suppression (also termed sense suppression) does not require that the entire transcription factor cDNA be introduced into the plant cells, nor does it require that the 20 introduced sequence be exactly identical to the endogenous transcription factor gene of interest. However, as with antisense suppression, the suppressive efficiency will be enhanced as specificity of hybridization is increased, e.g., as the introduced sequence is lengthened, and/or as the sequence similarity between the introduced sequence and the endogenous transcription factor gene is increased.

Vectors expressing an untranslatable form of the transcription factor mRNA, e.g., sequences comprising one or more stop codon, or nonsense mutation) can also be used to suppress expression of an endogenous transcription factor, thereby reducing or eliminating its activity and modifying one or more traits. Methods for producing such constructs are described in U.S. Patent No. 5,583,021. Preferably, such constructs are made by introducing 30 a premature stop codon into the transcription factor gene. Alternatively, a plant trait can be modified by gene silencing using double-strand RNA (Sharp (1999) Genes and Development 13: 139-141).

Another method for abolishing the expression of a gene is by insertion mutagenesis using the T-DNA of *Agrobacterium tumefaciens*. After generating the insertion 35 mutants, the mutants can be screened to identify those containing the insertion in a transcription factor or transcription factor homologue gene. Plants containing a single

transgene insertion event at the desired gene can be crossed to generate homozygous plants for the mutation (Koncz et al. (1992) Methods in Arabidopsis Research, World Scientific).

Alternatively, a plant phenotype can be altered by eliminating an endogenous gene, such as a transcription factor or transcription factor homologue, e.g., by homologous recombination (Kempin et al. (1997) Nature 389:802).

5 A plant trait can also be modified by using the cre-lox system (for example, as described in US Patent No. 5,658,772). A plant genome can be modified to include first and second lox sites that are then contacted with a Cre recombinase. If the lox sites are in the same orientation, the intervening DNA sequence between the two sites is excised. If the lox 10 sites are in the opposite orientation, the intervening sequence is inverted.

The polynucleotides and polypeptides of this invention can also be expressed in a plant in the absence of an expression cassette by manipulating the activity or expression level of the endogenous gene by other means. For example, by ectopically expressing a gene by T-DNA activation tagging (Ichikawa et al. (1997) Nature 390 698-701; Kakimoto et al. 15 (1996) Science 274: 982-985). This method entails transforming a plant with a gene tag containing multiple transcriptional enhancers and once the tag has inserted into the genome, expression of a flanking gene coding sequence becomes deregulated. In another example, the transcriptional machinery in a plant can be modified so as to increase transcription levels of a polynucleotide of the invention (See, e.g., PCT Publications WO 96/06166 and WO 98/53057 20 which describe the modification of the DNA binding specificity of zinc finger proteins by changing particular amino acids in the DNA binding motif).

The transgenic plant can also include the machinery necessary for expressing or altering the activity of a polypeptide encoded by an endogenous gene, for example by altering the phosphorylation state of the polypeptide to maintain it in an activated state.

25 Transgenic plants (or plant cells, or plant explants, or plant tissues) incorporating the polynucleotides of the invention and/or expressing the polypeptides of the invention can be produced by a variety of well established techniques as described above. Following construction of a vector, most typically an expression cassette, including a polynucleotide, e.g., encoding a transcription factor or transcription factor homologue, of the 30 invention, standard techniques can be used to introduce the polynucleotide into a plant, a plant cell, a plant explant or a plant tissue of interest. Optionally, the plant cell, explant or tissue can be regenerated to produce a transgenic plant.

The plant can be any higher plant, including gymnosperms, monocotyledonous and dicotyledonous plants. Suitable protocols are available for 35 *Leguminosae* (alfalfa, soybean, clover, etc.), *Umbelliferae* (carrot, celery, parsnip), *Cruciferae* (cabbage, radish, rapeseed, broccoli, etc.), *Cucurbitaceae* (melons and cucumber), *Gramineae* (wheat, corn, rice, barley, millet, etc.), *Solanaceae* (potato, tomato, tobacco,

peppers, etc.), and various other crops. See protocols described in Ammirato et al. (1984) Handbook of Plant Cell Culture -Crop Species. Macmillan Publ. Co. Shimamoto et al. (1989) Nature 338:274-276; Fromm et al. (1990) Bio/Technology 8:833-839; and Vasil et al. (1990) Bio/Technology 8:429-434.

5 Transformation and regeneration of both monocotyledonous and dicotyledonous plant cells is now routine, and the selection of the most appropriate transformation technique will be determined by the practitioner. The choice of method will vary with the type of plant to be transformed; those skilled in the art will recognize the suitability of particular methods for given plant types. Suitable methods can include, but are  
10 not limited to: electroporation of plant protoplasts; liposome-mediated transformation; polyethylene glycol (PEG) mediated transformation; transformation using viruses; micro-injection of plant cells; micro-projectile bombardment of plant cells; vacuum infiltration; and *Agrobacterium tumefaciens* mediated transformation. Transformation means introducing a nucleotide sequence in a plant in a manner to cause stable or transient expression of the  
15 sequence.

Successful examples of the modification of plant characteristics by transformation with cloned sequences which serve to illustrate the current knowledge in this field of technology, and which are herein incorporated by reference, include: U.S. Patent Nos. 5,571,706; 5,677,175; 5,510,471; 5,750,386; 5,597,945; 5,589,615; 5,750,871; 5,268,526;  
20 5,780,708; 5,538,880; 5,773,269; 5,736,369 and 5,610,042.

Following transformation, plants are preferably selected using a dominant selectable marker incorporated into the transformation vector. Typically, such a marker will confer antibiotic or herbicide resistance on the transformed plants, and selection of transformants can be accomplished by exposing the plants to appropriate concentrations of the  
25 antibiotic or herbicide.

After transformed plants are selected and grown to maturity, those plants showing a modified trait are identified. The modified trait can be any of those traits described above. Additionally, to confirm that the modified trait is due to changes in expression levels or activity of the polypeptide or polynucleotide of the invention can be determined by analyzing mRNA expression using Northern blots, RT-PCR or microarrays, or protein expression using immunoblots or Western blots or gel shift assays.

#### INTEGRATED SYSTEMS—SEQUENCE IDENTITY

Additionally, the present invention may be an integrated system, computer or computer readable medium that comprises an instruction set for determining the identity of  
35 one or more sequences in a database. In addition, the instruction set can be used to generate or identify sequences that meet any specified criteria. Furthermore, the instruction set may

be used to associate or link certain functional benefits, such improved pathogen tolerance, with one or more identified sequence.

For example, the instruction set can include, e.g., a sequence comparison or other alignment program, e.g., an available program such as, for example, the Wisconsin 5 Package Version 10.0, such as BLAST, FASTA, PILEUP, FINDPATTERNS or the like (GCG, Madison, WI). Public sequence databases such as GenBank, EMBL, Swiss-Prot and PIR or private sequence databases such as PhytoSeq (Incyte Pharmaceuticals, Palo Alto, CA) can be searched.

Alignment of sequences for comparison can be conducted by the local 10 homology algorithm of Smith and Waterman (1981) Adv. Appl. Math. 2:482, by the homology alignment algorithm of Needleman and Wunsch (1970) J. Mol. Biol. 48:443, by the search for similarity method of Pearson and Lipman (1988) Proc. Natl. Acad. Sci. U.S.A. 85: 15 2444, by computerized implementations of these algorithms. After alignment, sequence comparisons between two (or more) polynucleotides or polypeptides are typically performed by comparing sequences of the two sequences over a comparison window to identify and compare local regions of sequence similarity. The comparison window can be a segment of at least about 20 contiguous positions, usually about 50 to about 200, more usually about 100 to about 150 contiguous positions. A description of the method is provided in Ausubel et al., 20 *supra*.

A variety of methods of determining sequence relationships can be used, including manual alignment and computer assisted sequence alignment and analysis. This later approach is a preferred approach in the present invention, due to the increased throughput afforded by computer assisted methods. As noted above, a variety of computer programs for performing sequence alignment are available, or can be produced by one of 25 skill.

One example algorithm that is suitable for determining percent sequence identity and sequence similarity is the BLAST algorithm, which is described in Altschul et al. J. Mol. Biol. 215:403-410 (1990). Software for performing BLAST analyses is publicly available, e.g., through the National Center for Biotechnology Information 30 (<http://www.ncbi.nlm.nih.gov/>). This algorithm involves first identifying high scoring sequence pairs (HSPs) by identifying short words of length W in the query sequence, which either match or satisfy some positive-valued threshold score T when aligned with a word of the same length in a database sequence. T is referred to as the neighborhood word score threshold (Altschul et al., *supra*). These initial neighborhood word hits act as seeds for 35 initiating searches to find longer HSPs containing them. The word hits are then extended in both directions along each sequence for as far as the cumulative alignment score can be increased. Cumulative scores are calculated using, for nucleotide sequences, the parameters

M (reward score for a pair of matching residues; always  $> 0$ ) and N (penalty score for mismatching residues; always  $< 0$ ). For amino acid sequences, a scoring matrix is used to calculate the cumulative score. Extension of the word hits in each direction are halted when:

5 the cumulative alignment score falls off by the quantity X from its maximum achieved value;

10 the cumulative score goes to zero or below, due to the accumulation of one or more negative-scoring residue alignments; or the end of either sequence is reached. The BLAST algorithm parameters W, T, and X determine the sensitivity and speed of the alignment. The BLASTN program (for nucleotide sequences) uses as defaults a wordlength (W) of 11, an expectation (E) of 10, a cutoff of 100, M=5, N=-4, and a comparison of both strands. For amino acid

15 sequences, the BLASTP program uses as defaults a wordlength (W) of 3, an expectation (E) of 10, and the BLOSUM62 scoring matrix (*see* Henikoff & Henikoff (1989) Proc. Natl. Acad. Sci. USA 89:10915).

In addition to calculating percent sequence identity, the BLAST algorithm also performs a statistical analysis of the similarity between two sequences (*see*, e.g., Karlin & Altschul (1993) Proc. Natl. Acad. Sci. USA 90:5873-5787). One measure of similarity provided by the BLAST algorithm is the smallest sum probability (P(N)), which provides an indication of the probability by which a match between two nucleotide or amino acid sequences would occur by chance. For example, a nucleic acid is considered similar to a reference sequence (and, therefore, in this context, homologous) if the smallest sum probability in a comparison of the test nucleic acid to the reference nucleic acid is less than about 0.1, or less than about 0.01, and or even less than about 0.001. An additional example of a useful sequence alignment algorithm is PILEUP. PILEUP creates a multiple sequence alignment from a group of related sequences using progressive, pairwise alignments. The program can align, e.g., up to 300 sequences of a maximum length of 5,000 letters.

25 The integrated system, or computer typically includes a user input interface allowing a user to selectively view one or more sequence records corresponding to the one or more character strings, as well as an instruction set which aligns the one or more character strings with each other or with an additional character string to identify one or more region of sequence similarity. The system may include a link of one or more character strings with a particular phenotype or gene function. Typically, the system includes a user readable output element which displays an alignment produced by the alignment instruction set.

The methods of this invention can be implemented in a localized or distributed computing environment. In a distributed environment, the methods may be implemented on a single computer comprising multiple processors or on a multiplicity of computers. The computers can be linked, e.g. through a common bus, but more preferably the computer(s) are nodes on a network. The network can be a generalized or a dedicated local or

wide-area network and, in certain preferred embodiments, the computers may be components of an intra-net or an internet.

Thus, the invention provides methods for identifying a sequence similar or homologous to one or more polynucleotides as noted herein, or one or more target polypeptides encoded by the polynucleotides, or otherwise noted herein and may include linking or associating a given plant phenotype or gene function with a sequence. In the methods, a sequence database is provided (locally or across an inter or intra net) and a query is made against the sequence database using the relevant sequences herein and associated plant phenotypes or gene functions.

Any sequence herein can be entered into the database, before or after querying the database. This provides for both expansion of the database and, if done before the querying step, for insertion of control sequences into the database. The control sequences can be detected by the query to ensure the general integrity of both the database and the query. As noted, the query can be performed using a web browser based interface. For example, the database can be a centralized public database such as those noted herein, and the querying can be done from a remote terminal or computer across an internet or intranet.

#### EXAMPLES

The following examples are intended to illustrate but not limit the present invention.

20 **EXAMPLE I. FULL LENGTH GENE IDENTIFICATION AND CLONING**

Putative transcription factor sequences (genomic or ESTs) related to known transcription factors were identified in the *Arabidopsis thaliana* GenBank database using the tblastn sequence analysis program using default parameters and a P-value cutoff threshold of -4 or -5 or lower, depending on the length of the query sequence. Putative transcription factor sequence hits were then screened to identify those containing particular sequence strings. If the sequence hits contained such sequence strings, the sequences were confirmed as transcription factors.

Alternatively, *Arabidopsis thaliana* cDNA libraries derived from different tissues or treatments, or genomic libraries were screened to identify novel members of a transcription family using a low stringency hybridization approach. Probes were synthesized using gene specific primers in a standard PCR reaction (annealing temperature 60° C) and labeled with <sup>32</sup>P dCTP using the High Prime DNA Labeling Kit (Boehringer Mannheim). Purified radiolabelled probes were added to filters immersed in Church hybridization medium (0.5 M NaPO<sub>4</sub> pH 7.0, 7% SDS, 1 % w/v bovine serum albumin) and hybridized overnight at 60 °C with shaking. Filters were washed two times for 45 to 60 minutes with 1xSCC, 1% SDS at 60° C.

To identify additional sequence 5' or 3' of a partial cDNA sequence in a cDNA library, 5' and 3' rapid amplification of cDNA ends (RACE) was performed using the Marathon™ cDNA amplification kit (Clontech, Palo Alto, CA). Generally, the method entailed first isolating poly(A) mRNA, performing first and second strand cDNA synthesis to generate double stranded cDNA, blunting cDNA ends, followed by ligation of the Marathon™ Adaptor to the cDNA to form a library of adaptor-ligated ds cDNA.

Gene-specific primers were designed to be used along with adaptor specific primers for both 5' and 3' RACE reactions. Nested primers, rather than single primers, were used to increase PCR specificity. Using 5' and 3' RACE reactions, 5' and 3' RACE fragments were obtained, sequenced and cloned. The process can be repeated until 5' and 3' ends of the full-length gene were identified. Then the full-length cDNA was generated by PCR using primers specific to 5' and 3' ends of the gene by end-to-end PCR.

#### EXAMPLE II. CONSTRUCTION OF EXPRESSION VECTORS

The sequence was amplified from a genomic or cDNA library using primers specific to sequences upstream and downstream of the coding region. The expression vector was pMEN20 or pMEN65, which are both derived from pMON316 (Sanders et al, (1987) *Nucleic Acids Research* 15:1543-58) and contain the CaMV 35S promoter to express transgenes. To clone the sequence into the vector, both pMEN20 and the amplified DNA fragment were digested separately with SalI and NotI restriction enzymes at 37° C for 2 hours. The digestion products were subject to electrophoresis in a 0.8% agarose gel and visualized by ethidium bromide staining. The DNA fragments containing the sequence and the linearized plasmid were excised and purified by using a Qiaquick gel extraction kit (Qiagen, CA). The fragments of interest were ligated at a ratio of 3:1 (vector to insert). Ligation reactions using T4 DNA ligase (New England Biolabs, MA) were carried out at 16° C for 16 hours. The ligated DNAs were transformed into competent cells of the *E. coli* strain DH5alpha by using the heat shock method. The transformations were plated on LB plates containing 50 mg/l kanamycin (Sigma).

Individual colonies were grown overnight in five milliliters of LB broth containing 50 mg/l kanamycin at 37° C. Plasmid DNA was purified by using Qiaquick Mini Prep kits (Qiagen, CA).

#### EXAMPLE III. TRANSFORMATION OF AGROBACTERIUM WITH THE EXPRESSION VECTOR

After the plasmid vector containing the gene was constructed, the vector was used to transform *Agrobacterium tumefaciens* cells expressing the gene products. The stock of *Agrobacterium tumefaciens* cells for transformation were made as described by Nagel et al. (1990) *FEMS Microbiol Letts.* 67: 325-328. *Agrobacterium* strain ABI was grown in 250 ml

LB medium (Sigma) overnight at 28°C with shaking until an absorbance ( $A_{600}$ ) of 0.5 – 1.0 was reached. Cells were harvested by centrifugation at 4,000  $\times$  g for 15 min at 4° C. Cells were then resuspended in 250  $\mu$ l chilled buffer (1 mM HEPES, pH adjusted to 7.0 with KOH). Cells were centrifuged again as described above and resuspended in 125  $\mu$ l chilled buffer. Cells were then centrifuged and resuspended two more times in the same HEPES buffer as described above at a volume of 100  $\mu$ l and 750  $\mu$ l, respectively. Resuspended cells were then distributed into 40  $\mu$ l aliquots, quickly frozen in liquid nitrogen, and stored at -80° C.

*Agrobacterium* cells were transformed with plasmids prepared as described above following the protocol described by Nagel et al. For each DNA construct to be transformed, 50 – 100 ng DNA (generally resuspended in 10 mM Tris-HCl, 1 mM EDTA, pH 8.0) was mixed with 40  $\mu$ l of *Agrobacterium* cells. The DNA/cell mixture was then transferred to a chilled cuvette with a 2mm electrode gap and subject to a 2.5 kV charge dissipated at 25  $\mu$ F and 200  $\mu$ F using a Gene Pulser II apparatus (Bio-Rad). After electroporation, cells were immediately resuspended in 1.0 ml LB and allowed to recover without antibiotic selection for 2 – 4 hours at 28° C in a shaking incubator. After recovery, cells were plated onto selective medium of LB broth containing 100  $\mu$ g/ml spectinomycin (Sigma) and incubated for 24-48 hours at 28° C. Single colonies were then picked and inoculated in fresh medium. The presence of the plasmid construct was verified by PCR amplification and sequence analysis.

EXAMPLE IV. TRANSFORMATION OF *ARABIDOPSIS* PLANTS WITH *AGROBACTERIUM TUMEFACIENS* WITH EXPRESSION VECTOR

After transformation of *Agrobacterium tumefaciens* with plasmid vectors containing the gene, single *Agrobacterium* colonies were identified, propagated, and used to transform *Arabidopsis* plants. Briefly, 500 ml cultures of LB medium containing 50 mg/l kanamycin were inoculated with the colonies and grown at 28° C with shaking for 2 days until an absorbance ( $A_{600}$ ) of > 2.0 is reached. Cells were then harvested by centrifugation at 4,000  $\times$  g for 10 min, and resuspended in infiltration medium (1/2 X Murashige and Skoog salts (Sigma), 1 X Gamborg's B-5 vitamins (Sigma), 5.0% (w/v) sucrose (Sigma), 0.044  $\mu$ M benzylamino purine (Sigma), 200  $\mu$ l/L Silwet L-77 (Lehle Seeds) until an absorbance ( $A_{600}$ ) of 0.8 was reached.

Prior to transformation, *Arabidopsis thaliana* seeds (ecotype Columbia) were sown at a density of ~10 plants per 4" pot onto Pro-Mix BX potting medium (Hummert International) covered with fiberglass mesh (18 mm X 16 mm). Plants were grown under continuous illumination (50-75  $\mu$ E/m<sup>2</sup>/sec) at 22-23° C with 65-70% relative humidity. After about 4 weeks, primary inflorescence stems (bolts) are cut off to encourage growth of

multiple secondary bolts. After flowering of the mature secondary bolts, plants were prepared for transformation by removal of all siliques and opened flowers.

The pots were then immersed upside down in the mixture of *Agrobacterium* infiltration medium as described above for 30 sec, and placed on their sides to allow draining  
5 into a 1' x 2' flat surface covered with plastic wrap. After 24 h, the plastic wrap was removed and pots are turned upright. The immersion procedure was repeated one week later, for a total of two immersions per pot. Seeds were then collected from each transformation pot and analyzed following the protocol described below.

EXAMPLE V. IDENTIFICATION OF ARABIDOPSIS PRIMARY TRANSFORMANTS

10 Seeds collected from the transformation pots were sterilized essentially as follows. Seeds were dispersed into in a solution containing 0.1% (v/v) Triton X-100 (Sigma) and sterile H<sub>2</sub>O and washed by shaking the suspension for 20 min. The wash solution was then drained and replaced with fresh wash solution to wash the seeds for 20 min with shaking. After removal of the second wash solution, a solution containing 0.1% (v/v) Triton X-100 and  
15 70% ethanol (Equistar) was added to the seeds and the suspension was shaken for 5 min. After removal of the ethanol/detergent solution, a solution containing 0.1% (v/v) Triton X-100 and 30% (v/v) bleach (Clorox) was added to the seeds, and the suspension was shaken for 10 min. After removal of the bleach/detergent solution, seeds were then washed five times in sterile distilled H<sub>2</sub>O. The seeds were stored in the last wash water at 4° C for 2 days in the  
20 dark before being plated onto antibiotic selection medium (1 X Murashige and Skoog salts (pH adjusted to 5.7 with 1M KOH), 1 X Gamborg's B-5 vitamins, 0.9% phytagar (Life Technologies), and 50 mg/l kanamycin). Seeds were germinated under continuous illumination (50-75 μE/m<sup>2</sup>/sec) at 22-23° C. After 7-10 days of growth under these conditions, kanamycin resistant primary transformants (T<sub>1</sub> generation) were visible and  
25 obtained. These seedlings were transferred first to fresh selection plates where the seedlings continued to grow for 3-5 more days, and then to soil (Pro-Mix BX potting medium).

Primary transformants were crossed and progeny seeds (T<sub>2</sub>) collected; kanamycin resistant seedlings were selected and analyzed. The expression levels of the recombinant polynucleotides in the transformants varies from about a 5% expression level  
30 increase to a least a 100% expression level increase. Similar observations are made with respect to polypeptide level expression.

EXAMPLE VI. IDENTIFICATION OF ARABIDOPSIS PLANTS WITH TRANSCRIPTION FACTOR GENE KNOCKOUTS

The screening of insertion mutagenized *Arabidopsis* collections for null mutants in a known target gene was essentially as described in Krysan et al (1999) *Plant Cell* 11:2283-2290. Briefly, gene-specific primers, nested by 5-250 bases to each other, were designed from the 5' and 3' regions of a known target gene. Similarly, nested sets of primers 5 were also created specific to each of the T-DNA or transposon ends (the "right" and "left" borders). All possible combinations of gene specific and T-DNA/transposon primers were used to detect by PCR an insertion event within or close to the target gene. The amplified DNA fragments were then sequenced which allows the precise determination of the T-DNA/transposon insertion point relative to the target gene. Insertion events within the coding 10 or intervening sequence of the genes were deconvoluted from a pool comprising a plurality of insertion events to a single unique mutant plant for functional characterization. The method is described in more detail in Yu and Adam, US Application Serial No. 09/177,733 filed October 23, 1998.

15 **EXAMPLE VII. IDENTIFICATION OF PATHOGEN INDUCED GENES**

In some instances, expression patterns of the pathogen induced genes (such as defense genes) was monitored by microarray experiments. cDNAs were generated by PCR and resuspended at a final concentration of ~ 100 ng/ul in 3X SSC or 150mM Na-phosphate (Eisen and Brown (1999) *Meth. in Enzymol.* 303:179-205). The cDNAs were spotted on 20 microscope glass slides coated with polylysine. The prepared cDNAs were aliquoted into 384 well plates and spotted on the slides using an x-y-z gantry (OmniGrid) purchased from GeneMachines (Menlo Park, CA) outfitted with quill type pins purchased from Telechem International (Sunnyvale, CA). After spotting, the arrays were cured for a minimum of one week at room temperature, rehydrated and blocked following the protocol recommended by 25 Eisen and Brown (1999).

Sample total RNA (10 ug) samples were labeled using fluorescent Cy3 and Cy5 dyes. Labeled samples were resuspended in 4X SSC/0.03% SDS/4 ug salmon sperm DNA/2 ug tRNA/ 50mM Na-pyrophosphate, heated for 95°C for 2.5 minutes, spun down and placed on the array. The array was then covered with a glass coverslip and placed in a sealed chamber. 30 The chamber was then kept in a water bath at 62°C overnight. The arrays were washed as described in Eisen and Brown (1999) and scanned on a General Scanning 3000 laser scanner. The resulting files are subsequently quantified using Imagene a software purchased from BioDiscovery (Los Angeles, CA).

EXAMPLE VIII. IDENTIFICATION OF PATHOGEN TOLERANCE PHENOTYPE IN OVEREXPRESSOR OR GENE KNOCKOUT PLANTS

Experiments were performed to identify those transformants or knockouts that exhibited an improved pathogen tolerance. For such studies, the transformants were exposed to biotrophic fungal pathogens, such as *Erysiphe orontii*; and necrotrophic fungal pathogens, such as *Fusarium oxysporum*. *Fusarium oxysporum* isolates cause vascular wilts and damping off of various annual vegetables, perennials and weeds (Mauch-Mani and Slusarenko (1994) Molecular Plant-Microbe Interactions 7: 378-383). For *Fusarium oxysporum* experiments, plants grown on petri dishes were sprayed with a fresh spore suspension of *F. oxysporum*. The spore suspension was prepared as follows: A plug of fungal hyphae from a plate culture was placed on a fresh potato dextrose agar plate and allowed to spread for one week. 5 ml sterile water was then added to the plate, swirled, and pipetted into 50 ml Armstrong Fusarium medium. Spores were grown overnight in Fusarium medium and then sprayed onto plants using a Preval paint sprayer. Plant tissue was harvested and frozen in liquid nitrogen 48 hours post infection.

*Erysiphe orontii* is a causal agent of powdery mildew. For *Erysiphe orontii* experiments, plants were grown approximately 4 weeks in a greenhouse under 12 hour light (20 C, ~30% relative humidity (rh)). Individual leaves were infected with *E. orontii* spores from infected plants using a camel's hair brush, and the plants were transferred to a Percival growth chamber (20 C, 80% rh.). Plant tissue was harvested and frozen in liquid nitrogen 7 days post infection.

*Botrytis cinerea* is a necrotrophic pathogen. *Botrytis cinerea* was grown on potato dextrose agar in the light. A spore culture was made by spreading 10 ml of sterile water on the fungus plate, swirling and transferring spores to 10 ml of sterile water. The spore inoculum (approx. 105 spores/ml) was used to spray 10 day-old seedlings grown under sterile conditions on MS (-sucrose) media. Symptoms were evaluated every day up to approximately 1 week.

Infection with bacterial pathogens *Pseudomonas syringae* pv *maculicola* strain 4326 and pv *maculicola* strain 4326 was performed by hand inoculation at two doses. Two inoculation doses allows the differentiation between plants with enhanced susceptibility and plants with enhanced resistance to the pathogen. Plants were grown for 3 weeks in the greenhouse, then transferred to the growth chamber for the remainder of their growth. Psm ES4326 was hand inoculated with 1 ml syringe on 3 fully-expanded leaves per plant (4 1/2 wk old), using at least 9 plants per overexpressing line at two inoculation doses, OD=0.005 and OD=0.0005. Disease scoring occurred at day 3 post-inoculation with pictures of the plants and leaves taken in parallel.

Table 3 shows the phenotypes observed for particular overexpressor or knockout plants and provides the SEQ ID No., the internal reference code (GID), whether a knockout or overexpressor plant was analyzed and the observed phenotype.

**Table 3**

SEQ ID No.	GID	Knockout (KO) or overexpressor (OE)	Phenotype
1	G188	KO	Increased susceptibility to Fusarium
3	G616	OE	Increased tolerance to Erysiphe
5	G19	OE	Increased tolerance to Erysiphe
7	G261	OE	Increased susceptibility to Botrytis
9	G28	OE	Increased resistance to Erysiphe
11	G869	OE	Increased susceptibility to Fusarium
13	G237	OE	Increased tolerance to Erysiphe
15	G409	OE	Increased tolerance to Erysiphe
17	G418	OE	Increased tolerance to Pseudomonas
19	G591	OE	Increased tolerance to Erysiphe
21	G525	OE	Increased tolerance to Pseudomonas
23	G545	OE	Increased susceptibility to Pseudomonas, Erysiphe and Fusarium
25	G865	OE	Increased susceptibility to Erysiphe and Botrytis
27	G881	OE	Increased susceptibility to Erysiphe and Botrytis
29	G896	KO	Increased susceptibility to Fusarium
31	G378	OE	Increased resistance to Erysiphe
33	G569	OE	Decreased expression of defense genes
35	G558	OE	Increased expression of defense genes

5 For a particular overexpressor that shows an increased susceptibility to a pathogen, it may be more useful to select a plant with a decreased expression of the particular transcription factor. For a particular knockout that shows an increased susceptibility to a pathogen, it may be more useful to select a plant with an increased expression of the particular transcription factor.

10 Other than *Fusarium oxysporum*, *Erysiphe orontii*, the transgenic plants are more tolerant to *Sclerotinia spp.*, soil-borne oomycetes, foliar oomycetes, *Botrytis spp.*, *Rhizoctonia spp.*, *Verticillium dahliae/albo-atrum*, *Alternaria spp.*, rusts, *Mycosphaerella spp.*, *Fusarium solani*, or the like. The transgenic plants are more resistant to fungal diseases such as rusts, smuts, wilts, yellows, root rot, leaf drop, ergot, leaf blight of potato, brown spot of rice, leaf blight, late blight, powdery mildew, downy mildew, and the like; viral diseases such as sugarcane mosaic, cassava mosaic, sugar beet yellows, plum pox, barley yellow dwarf, tomato yellow leaf curl, tomato spotted wilt virus, and the like; bacterial diseases such as citrus canker, bacterial leaf blight, bacterial wilt, soft rot of vegetables, and the like; nematode diseases such as root knot, sugar beet cyst nematode or the like.

EXAMPLE IX. IDENTIFICATION OF HOMOLOGOUS SEQUENCES

Homologous sequences from *Arabidopsis* and plant species other than *Arabidopsis* were identified using database sequence search tools, such as the Basic Local Alignment Search Tool (BLAST) (Altschul et al. (1990) *J. Mol. Biol.* 215:403-410; and Altschul et al.

5 (1997) *Nucl. Acid Res.* 25: 3389-3402). The tblastx sequence analysis programs were employed using the BLOSUM-62 scoring matrix (Henikoff, S. and Henikoff, J. G. (1992) *Proc. Natl. Acad. Sci. USA* 89: 10915-10919).

Identified *Arabidopsis* homologous sequences are provided in Figure 2 and included in the Sequence Listing. The percent sequence identity among these sequences is as low as 10 47% sequence identity. Additionally, the entire NCBI GenBank database was filtered for sequences from all plants except *Arabidopsis thaliana* by selecting all entries in the NCBI GenBank database associated with NCBI taxonomic ID 33090 (Viridiplantae; all plants) and excluding entries associated with taxonomic ID 3701 (*Arabidopsis thaliana*). These sequences were compared to sequences representing genes of SEQ IDs Nos. 1-58 on 15 9/26/2000 using the Washington University TBLASTX algorithm (version 2.0a19MP). For each gene of SEQ IDs Nos. 1-58, individual comparisons were ordered by probability score (P-value), where the score reflects the probability that a particular alignment occurred by chance. For example, a score of 3.6e-40 is  $3.6 \times 10^{-40}$ . For up to ten species, the gene with the lowest P-value (and therefore the most likely homolog) is listed in Figure 3.

20 In addition to P-values, comparisons were also scored by percentage identity. Percentage identity reflects the degree to which two segments of DNA or protein are identical over a particular length. The ranges of percent identity between the non-*Arabidopsis* genes shown in Figure 3 and the *Arabidopsis* genes in the sequence listing are: SEQ ID No. 1: 38%-76%; SEQ ID No. 3: 36%-72%; SEQ ID No. 5: 51%-75%; SEQ ID No. 7: 37%-76%; SEQ ID 25 No. 9: 48%-75%; SEQ ID No. 11: 31%-68%; SEQ ID No. 13: 59%-81%; SEQ ID No. 15: 49%-81%; SEQ ID No. 17: 53%-87%; SEQ ID No. 19: 48%-84%; SEQ ID No. 21: 73%-89%; SEQ ID No. 23: 52%-64%; SEQ ID No. 25: 48%-83%; SEQ ID No. 27: 35%-92%; SEQ ID No. 29: 56%-89%; SEQ ID No. 31: 50%-90%; SEQ ID No. 33: 50%-93%; SEQ ID No. 35: 52%-81%; SEQ ID No. 37: 75%-81%; SEQ ID No. 39: 35%-72%; SEQ ID No. 41: 55%-89%; SEQ ID No. 43: 56%-77%; SEQ ID No. 45: 34%-72%; SEQ ID No. 47: 51%-86%; SEQ ID No. 49: 46%-86%; SEQ ID No. 51: 58%-80%; SEQ ID No. 53: 46%-55%; SEQ ID No. 55: 84%-89%; and SEQ ID No. 57: 43%-71%.

The polynucleotides and polypeptides in the Sequence Listing and the identified homologous sequences may be stored in a computer system and have associated or linked 35 with the sequences a function, such as that the polynucleotides and polypeptides are useful for modifying the pathogen tolerance of a plant.

All references, publications, patents and other documents herein are incorporated by reference in their entirety for all purposes. Although the invention has been described with reference to the embodiments and examples above, it should be understood that various modifications can be made without departing from the spirit of the invention.

What is claimed is:

1. A transgenic plant with modified pathogen tolerance, which plant comprises a recombinant polynucleotide comprising a nucleotide sequence selected from the group consisting of:
  - 5 (a) a nucleotide sequence encoding a polypeptide comprising a sequence selected from SEQ ID Nos. 2N, where N=1-29, or a complementary nucleotide sequence thereof;
  - (b) a nucleotide sequence encoding a polypeptide comprising a conservatively substituted variant of a polypeptide of (a);
  - 10 (c) a nucleotide sequence comprising a sequence selected from those of SEQ ID Nos. 2N-1, where N=1-29, or a complementary nucleotide sequence thereof;
  - (d) a nucleotide sequence comprising silent substitutions in a nucleotide sequence of (c);
  - (e) a nucleotide sequence which hybridizes under stringent conditions to a nucleotide sequence of one or more of: (a), (b), (c), or (d);
  - 15 (f) a nucleotide sequence comprising at least 15 consecutive nucleotides of a sequence of any of (a)-(e);
  - (g) a nucleotide sequence comprising a subsequence or fragment of any of (a)-(f), which subsequence or fragment encodes a polypeptide that modifies a plant's pathogen tolerance;
  - 20 (h) a nucleotide sequence having at least 31% sequence identity to a nucleotide sequence of any of (a)-(g);
  - (i) a nucleotide sequence having at least 60% identity sequence identity to a nucleotide sequence of any of (a)-(g);
  - 25 (j) a nucleotide sequence which encodes a polypeptide having at least 31% identity sequence identity to a polypeptide of SEQ ID Nos. 2N, where N=1-29;
  - (k) a nucleotide sequence which encodes a polypeptide having at least 60% identity sequence identity to a polypeptide of SEQ ID Nos. 2N, where N=1-29; and
  - 30 (l) a nucleotide sequence which encodes a polypeptide having at least 65% sequence identity to a conserved domain of a polypeptide of SEQ ID Nos. 2N, where N=1-29.
2. The transgenic plant of claim 1, further comprising a constitutive, inducible, or tissue-active promoter operably linked to said nucleotide sequence.
- 35 3. The transgenic plant of claim 1, wherein the plant is selected from the group consisting of: soybean, wheat, corn, potato, cotton, rice, oilseed rape, sunflower, alfalfa, sugarcane, turf, banana, blackberry, blueberry, strawberry, raspberry, cantaloupe, carrot,

cauliflower, coffee, cucumber, eggplant, grapes, honeydew, lettuce, mango, melon, onion, papaya, peas, peppers, pineapple, spinach, squash, sweet corn, tobacco, tomato, watermelon, rosaceous fruits, and vegetable brassicas.

- 5    4. An isolated or recombinant polynucleotide comprising a nucleotide sequence selected from the group consisting of:
- (a) a nucleotide sequence encoding a polypeptide comprising a sequence selected from SEQ ID Nos. 2N, where N=1-29, or a complementary nucleotide sequence thereof;
  - 10    (b) a nucleotide sequence encoding a polypeptide comprising a conservatively substituted variant of a polypeptide of (a);
  - (c) a nucleotide sequence comprising a sequence selected from those of SEQ ID Nos. 2N-1, where N=1-29, or a complementary nucleotide sequence thereof;
  - (d) a nucleotide sequence comprising silent substitutions in a nucleotide sequence of
  - 15    (c);
  - (e) a nucleotide sequence which hybridizes under stringent conditions to a nucleotide sequence of one or more of: (a), (b), (c), or (d);
  - (f) a nucleotide sequence comprising at least 15 consecutive nucleotides of a sequence of any of (a)-(e);
  - 20    (g) a nucleotide sequence comprising a subsequence or fragment of any of (a)-(f), which subsequence or fragment encodes a polypeptide having a biological activity that modifies a plant's pathogen tolerance;
  - (h) a nucleotide sequence having at least 31% sequence identity to a nucleotide sequence of any of (a)-(g);
  - 25    (i) a nucleotide sequence having at least 60% identity sequence identity to a nucleotide sequence of any of (a)-(g);
  - (j) a nucleotide sequence which encodes a polypeptide having at least 31% identity sequence identity to a polypeptide of SEQ ID Nos. 2N, where N=1-29;
  - (k) a nucleotide sequence which encodes a polypeptide having at least 60% identity sequence identity to a polypeptide of SEQ ID Nos. 2N, where N=1-29; and
  - 30    (l) a nucleotide sequence which encodes a conserved domain of a polypeptide having at least 65% sequence identity to a conserved domain of a polypeptide of SEQ ID Nos. 2N, where N=1-29.
- 35    5. The isolated or recombinant polynucleotide of claim 4, further comprising a constitutive, inducible, or tissue-active promoter operably linked to the nucleotide sequence.

6. A cloning or expression vector comprising the isolated or recombinant polynucleotide of claim 4.
7. A cell comprising the cloning or expression vector of claim 6.  
5
8. A transgenic plant comprising the isolated or recombinant polynucleotide of claim 4.
9. A composition produced by one or more of:
  - (a) incubating one or more polynucleotide of claim 4 with a nuclease;
  - 10 (b) incubating one or more polynucleotide of claim 4 with a restriction enzyme;
  - (c) incubating one or more polynucleotide of claim 4 with a polymerase;
  - (d) incubating one or more polynucleotide of claim 4 with a polymerase and a primer;
  - (e) incubating one or more polynucleotide of claim 4 with a cloning vector, or
  - (f) incubating one or more polynucleotide of claim 4 with a cell.
- 15
10. A composition comprising two or more different polynucleotides of claim 4.
11. An isolated or recombinant polypeptide comprising a subsequence of at least about 15 contiguous amino acids encoded by the recombinant or isolated polynucleotide of claim 4.  
20
12. A plant comprising an isolated polypeptide of claim 11.
13. A method for producing a plant having a modified pathogen tolerance, the method comprising altering the expression of the isolated or recombinant polynucleotide of claim 4 or  
25 the expression levels or activity of a polypeptide of claim 11 in a plant, thereby producing a modified plant, and selecting the modified plant for improved pathogen tolerance thereby providing the modified plant with a modified pathogen tolerance.
14. The method of claim 13, wherein the polynucleotide is a polynucleotide of claim 4.  
30
15. A method of identifying a factor that is modulated by or interacts with a polypeptide encoded by a polynucleotide of claim 4, the method comprising:
  - (a) expressing a polypeptide encoded by the polynucleotide in a plant; and
  - (b) identifying at least one factor that is modulated by or interacts with the  
35 polypeptide.

16. The method of claim 15, wherein the identifying is performed by detecting binding by the polypeptide to a promoter sequence, or detecting interactions between an additional protein and the polypeptide in a yeast two hybrid system.

5 17. The method of claim 15, wherein the identifying is performed by detecting expression of a factor by hybridization to a microarray, subtractive hybridization or differential display.

18. A method of identifying a molecule that modulates activity or expression of a polynucleotide or polypeptide of interest, the method comprising:

10 (a) placing the molecule in contact with a plant comprising the polynucleotide or polypeptide encoded by the polynucleotide of claim 4; and,

(b) monitoring one or more of:

(i) expression level of the polynucleotide in the plant;

(ii) expression level of the polypeptide in the plant;

15 (iii) modulation of an activity of the polypeptide in the plant; or

(iv) modulation of an activity of the polynucleotide in the plant.

19. An integrated system, computer or computer readable medium comprising one or more character strings corresponding to a polynucleotide of claim 4, or to a polypeptide encoded by the polynucleotide.

20 20. The integrated system, computer or computer readable medium of claim 19, further comprising a link between said one or more sequence strings to a modified plant pathogen tolerance phenotype.

25

21. A method of identifying a sequence similar or homologous to one or more polynucleotides of claim 4, or one or more polypeptides encoded by the polynucleotides, the method comprising:

(a) providing a sequence database; and,

30 (b) querying the sequence database with one or more target sequences corresponding to the one or more polynucleotides or to the one or more polypeptides to identify one or more sequence members of the database that display sequence similarity or homology to one or more of the one or more target sequences.

35 22. The method of claim 21, wherein the querying comprises aligning one or more of the target sequences with one or more of the one or more sequence members in the sequence database.

23. The method of claim 21, wherein the querying comprises identifying one or more of the one or more sequence members of the database that meet a user-selected identity criteria with one or more of the target sequences.

5

24. The method of claim 21, further comprising linking the one or more of the polynucleotides of claim 4, or encoded polypeptides, to a modified plant pathogen tolerance phenotype.

10 25. A plant comprising altered expression levels of an isolated or recombinant polynucleotide of claim 4.

26. A plant comprising altered expression levels or the activity of an isolated or recombinant polypeptide of claim 11.

15

27. A plant lacking a nucleotide sequence encoding a polypeptide of claim 11.

**Figure 1**

SEQ ID No.	GID	cDNA or protein	conserved domain
1	G188	cDNA	
2	G188	protein	175-222
3	G616	cDNA	
4	G616	protein	39-95
5	G19	cDNA	
6	G19	protein	76-145
7	G261	cDNA	
8	G261	protein	16-104
9	G28	cDNA	
10	G28	protein	145-213
11	G869	cDNA	
12	G869	protein	109-177
13	G237	cDNA	
14	G237	protein	11-113
15	G409	cDNA	
16	G409	protein	64-124
17	G418	cDNA	
18	G418	protein	500-560
19	G591	cDNA	
20	G591	protein	143-240
21	G525	cDNA	
22	G525	protein	23-167
23	G545	cDNA	
24	G545	protein	82-102, 136-154
25	G865	cDNA	
26	G865	protein	36-103
27	G881	cDNA	
28	G881	protein	176-233
29	G896	cDNA	
30	G896	protein	18-39
31	G378	cDNA	
32	G378	protein	196-237
33	G569	cDNA	
34	G569	protein	90-153
35	G558	cDNA	
36	G558	protein	45-105

**Figure 2**

SEQ ID No.	GID	homolog	cDNA or protein	conserved domain
37	G1396	homolog of G1394	cDNA	
38	G1396	homolog of G1394	protein	entire protein
39	G265	homolog of G261	cDNA	
40	G265	homolog of G261	protein	14-105
41	G1006	homolog of G28	cDNA	
42	G1006	homolog of G28	protein	114-182
43	G1309	homolog of G237	cDNA	
44	G1309	homolog of G237	protein	9-114
45	G2550	homolog of G418	cDNA	
46	G2550	homolog of G418	protein	348-408
47	G965	homolog of G418	cDNA	
48	G965	homolog of G418	protein	423-486
49	G793	homolog of G591	cDNA	
50	G793	homolog of G591	protein	151-206
51	G764	homolog of G525	cDNA	
52	G764	homolog of G525	protein	27-171
53	G350	homolog of G545	cDNA	
54	G350	homolog of G545	protein	91-113,150-170
55	G986	homolog of G881	cDNA	
56	G986	homolog of G881	protein	146-203
57	G1349	homolog of G896	cDNA	
58	G1349	homolog of G896	protein	13-63

Figure 3A

SEQ ID No.	GID	Genbank NID	P-value	Species
1	G188	7779802	5.20E-36	<i>Lotus japonicus</i>
1	G188	7284340	2.10E-34	<i>Glycine max</i>
1	G188	9361307	1.20E-27	<i>Triticum aestivum</i>
1	G188	7340336	1.10E-22	<i>Oryza sativa</i>
1	G188	6529152	3.60E-22	<i>Lycopersicon esculentum</i>
1	G188	8748477	7.70E-21	<i>Medicago truncatula</i>
1	G188	5456433	7.10E-14	<i>Zea mays</i>
1	G188	9302479	1.60E-12	<i>Sorghum bicolor</i>
1	G188	6696287	4.10E-12	<i>Pinus taeda</i>
1	G188	562242	9.00E-12	<i>Brassica rapa</i>
3	G616	7719440	8.30E-37	<i>Lotus japonicus</i>
3	G616	7692230	5.90E-33	<i>Glycine max</i>
3	G616	7501307	1.10E-21	<i>Gossypium arboreum</i>
3	G616	8071090	1.50E-21	<i>Solanum tuberosum</i>
3	G616	8858771	1.50E-21	<i>Oryza sativa</i>
3	G616	5047315	1.50E-21	<i>Gossypium hirsutum</i>
3	G616	6358532	5.80E-20	<i>Antirrhinum graniticum</i>
3	G616	2826867	7.00E-20	<i>Antirrhinum majus</i>
3	G616	6358535	7.40E-20	<i>Antirrhinum majus subsp. linkianum</i>
3	G616	6358538	7.50E-20	<i>Antirrhinum braun-blancetii</i>
5	G19	8789223	2.80E-34	<i>Citrus x paradisi</i>
5	G19	9434234	4.50E-34	<i>Lycopersicon esculentum</i>
5	G19	7478682	1.30E-30	<i>Glycine max</i>
5	G19	6654934	1.20E-28	<i>Medicago truncatula</i>
5	G19	3264766	5.50E-26	<i>Prunus armeniaca</i>
5	G19	7624302	8.30E-26	<i>Gossypium arboreum</i>
5	G19	9425363	2.90E-25	<i>Triticum aestivum</i>
5	G19	688579	3.60E-25	<i>Ricinus communis</i>
5	G19	9419304	6.00E-25	<i>Hordeum vulgare</i>
5	G19	7720316	8.80E-25	<i>Lotus japonicus</i>
7	G261	5821137	5.10E-93	<i>Nicotiana tabacum</i>
7	G261	7158881	8.80E-86	<i>Medicago sativa</i>
7	G261	886741	1.00E-73	<i>Zea mays</i>
7	G261	5900449	5.20E-47	<i>Lycopersicon esculentum</i>
7	G261	7561318	1.20E-46	<i>Medicago truncatula</i>
7	G261	19491	1.70E-42	<i>Lycopersicon peruvianum</i>
7	G261	7233914	3.50E-41	<i>Glycine max</i>
7	G261	4528238	9.00E-41	<i>Citrus unshiu</i>
7	G261	8903922	4.00E-39	<i>Hordeum vulgare</i>
7	G261	9251913	1.90E-36	<i>Solanum tuberosum</i>
9	G28	7528275	4.20E-62	<i>Mesembryanthemum crystallinum</i>
9	G28	6654776	1.20E-57	<i>Medicago truncatula</i>
9	G28	790362	2.30E-54	<i>Nicotiana tabacum</i>
9	G28	8809570	8.00E-54	<i>Nicotiana sylvestris</i>
9	G28	3342210	8.40E-54	<i>Lycopersicon esculentum</i>
9	G28	6566281	8.40E-47	<i>Glycine max</i>
9	G28	7627061	8.40E-47	<i>Gossypium arboreum</i>
9	G28	7324479	2.00E-44	<i>Lycopersicon pennellii</i>
9	G28	6478844	1.80E-35	<i>Matricaria chamomilla</i>
9	G28	7273972	7.80E-29	<i>Oryza sativa</i>
11	G869	2213784	1.30E-19	<i>Lycopersicon esculentum</i>
11	G869	3065894	7.30E-19	<i>Nicotiana tabacum</i>

Figure 3B

SEQ ID No.	GID	Genbank NID	P-value	Species
11	G869	8570080	4.20E-18	<i>Oryza sativa</i>
11	G869	7560260	1.50E-17	<i>Medicago truncatula</i>
11	G869	7534890	5.20E-14	<i>Sorghum bicolor</i>
11	G869	6455322	1.10E-13	<i>Glycine max</i>
11	G869	9362061	2.70E-13	<i>Triticum aestivum</i>
11	G869	7788764	5.70E-13	<i>Lotus japonicus</i>
11	G869	7624302	2.50E-12	<i>Gossypium arboreum</i>
11	G869	3858036	2.80E-12	<i>Populus balsamifera</i> subsp. <i>trichocarpa</i>
13	G237	8283916	4.70E-42	<i>Glycine max</i>
13	G237	9361969	8.30E-41	<i>Triticum aestivum</i>
13	G237	4753385	4.10E-39	<i>Zea mays</i>
13	G237	7535969	4.10E-33	<i>Sorghum bicolor</i>
13	G237	7566043	9.30E-33	<i>Medicago truncatula</i>
13	G237	7339127	2.00E-32	<i>Lycopersicon esculentum</i>
13	G237	5860031	1.10E-28	<i>Pinus taeda</i>
13	G237	7776223	2.20E-28	<i>Lotus japonicus</i>
13	G237	6850206	5.10E-28	<i>Oryza sativa</i>
13	G237	5048991	8.50E-28	<i>Gossypium hirsutum</i>
15	G409	6654773	6.10E-57	<i>Medicago truncatula</i>
15	G409	6531235	2.00E-56	<i>Lycopersicon esculentum</i>
15	G409	7924152	1.10E-47	<i>Glycine max</i>
15	G409	5006854	6.50E-43	<i>Oryza sativa</i>
15	G409	8098529	2.10E-41	<i>Hordeum vulgare</i>
15	G409	767697	1.40E-37	<i>Daucus carota</i>
15	G409	8328991	3.30E-37	<i>Mesembryanthemum crystallinum</i>
15	G409	7415613	1.40E-32	<i>Physcomitrella patens</i>
15	G409	7785121	2.80E-32	<i>Lotus japonicus</i>
15	G409	6916941	4.80E-32	<i>Lycopersicon pennellii</i>
17	G418	7239156	1.90E-123	<i>Malus x domestica</i>
17	G418	5892190	2.00E-62	<i>Lycopersicon esculentum</i>
17	G418	7628137	8.70E-58	<i>Gossypium arboreum</i>
17	G418	9205496	3.90E-51	<i>Glycine max</i>
17	G418	6069643	1.50E-45	<i>Oryza sativa</i>
17	G418	7562931	6.90E-45	<i>Medicago truncatula</i>
17	G418	7781695	5.50E-40	<i>Lotus japonicus</i>
17	G418	9298824	7.80E-34	<i>Sorghum bicolor</i>
17	G418	9428023	3.90E-32	<i>Triticum aestivum</i>
17	G418	7244366	1.30E-31	<i>Mentha x piperita</i>
19	G591	7646333	1.90E-55	<i>Lycopersicon esculentum</i>
19	G591	7924288	4.10E-53	<i>Glycine max</i>
19	G591	7722838	1.10E-41	<i>Lotus japonicus</i>
19	G591	5804781	1.40E-24	<i>Nicotiana tabacum</i>
19	G591	9198126	2.50E-23	<i>Medicago truncatula</i>
19	G591	427677	9.50E-15	<i>Oryza sativa</i>
19	G591	7624745	1.80E-14	<i>Gossypium arboreum</i>
19	G591	7535578	8.70E-14	<i>Sorghum bicolor</i>
19	G591	5915205	1.30E-11	<i>Zea mays</i>
19	G591	9249806	2.60E-11	<i>Solanum tuberosum</i>
21	G525	4384535	5.60E-61	<i>Lycopersicon esculentum</i>
21	G525	6454868	2.00E-58	<i>Glycine max</i>
21	G525	6066594	9.30E-54	<i>Petunia x hybrida</i>
21	G525	4977542	8.60E-51	<i>Oryza sativa</i>

Figure 3C

SEQ ID No.	GID	Genbank NID	P-value	Species
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21	G525	4218536	5.20E-50	Triticum sp.
21	G525	6732159	5.20E-50	Triticum monococcum
21	G525	5343151	2.70E-49	Zea mays
21	G525	5049217	4.20E-48	Gossypium hirsutum
21	G525	8708684	8.90E-48	Hordeum vulgare
23	G545	4666359	8.30E-55	Datisca glomerata
23	G545	7228328	3.70E-52	Medicago sativa
23	G545	1763062	1.30E-51	Glycine max
23	G545	7206360	3.10E-44	Medicago truncatula
23	G545	7626808	9.60E-40	Gossypium arboreum
23	G545	439492	3.90E-39	Petunia x hybrida
23	G545	4382658	1.70E-38	Lycopersicon esculentum
23	G545	8486215	8.70E-38	Euphorbia esula
23	G545	7322653	6.80E-37	Lycopersicon hirsutum
23	G545	7785845	1.10E-33	Lotus japonicus
25	G865	9417297	1.70E-32	Triticum aestivum
25	G865	7206394	4.90E-29	Medicago truncatula
25	G865	7796858	5.70E-27	Glycine max
25	G865	4387560	9.20E-25	Lycopersicon esculentum
25	G865	569065	1.50E-23	Oryza sativa
25	G865	7788764	4.10E-23	Lotus japonicus
25	G865	790362	8.40E-22	Nicotiana tabacum
25	G865	7528275	5.90E-21	Mesembryanthemum crystallinum
25	G865	3264766	8.80E-20	Prunus armeniaca
25	G865	8098026	2.00E-19	Hordeum vulgare
27	G881	5820418	9.80E-29	Glycine max
27	G881	8440065	1.00E-27	Gossypium hirsutum
27	G881	4380578	1.50E-27	Lycopersicon esculentum
27	G881	9199620	2.70E-27	Medicago truncatula
27	G881	6472584	2.20E-24	Nicotiana tabacum
27	G881	9250698	3.20E-24	Solanum tuberosum
27	G881	8205146	5.20E-21	Oryza sativa
27	G881	1159878	8.20E-17	Avena fatua
27	G881	9299778	2.70E-16	Sorghum bicolor
27	G881	9444636	1.10E-14	Triticum aestivum
29	G896	9410462	1.90E-101	Hordeum vulgare
29	G896	7628908	3.60E-82	Gossypium arboreum
29	G896	7244408	1.80E-79	Mentha x piperita
29	G896	5046180	2.10E-73	Gossypium hirsutum
29	G896	7678652	1.10E-63	Lotus japonicus
29	G896	8286031	1.40E-60	Glycine max
29	G896	5888938	4.50E-58	Lycopersicon esculentum
29	G896	9298238	9.20E-54	Sorghum bicolor
29	G896	7566414	8.00E-52	Medicago truncatula
29	G896	8845076	1.00E-46	Triticum aestivum
31	G378	5270028	5.10E-73	Lycopersicon esculentum
31	G378	5048335	4.10E-58	Gossypium hirsutum
31	G378	7239521	5.90E-42	Oryza sativa
31	G378	5606120	6.80E-36	Glycine max
31	G378	3853800	3.20E-30	Populus tremula x Populus tremuloides
31	G378	7659983	1.70E-23	Sorghum bicolor

Figure 3D

SEQ ID No.	GID	Genbank NID	P-value	Species
31	G378	6626305	1.10E-21	<i>Zea mays</i>
31	G378	9412941	9.40E-19	<i>Triticum aestivum</i>
31	G378	3242033	4.30E-17	<i>Mesembryanthemum crystallinum</i>
31	G378	7626259	7.70E-13	<i>Gossypium arboreum</i>
33	G229	7337390	6.60E-51	<i>Lycopersicon esculentum</i>
33	G229	9823237	3.60E-50	<i>Hordeum vulgare</i>
33	G229	7244424	4.90E-50	<i>Mentha x piperita</i>
33	G229	7776053	1.70E-49	<i>Lotus japonicus</i>
33	G229	2921335	5.80E-48	<i>Gossypium hirsutum</i>
33	G229	1491932	4.50E-47	<i>Zea mays</i>
33	G229	6455590	2.80E-44	<i>Glycine max</i>
33	G229	6020191	2.00E-41	<i>Pinus taeda</i>
33	G229	10697236	4.20E-41	<i>Oryza sativa</i>
33	G229	7765706	5.10E-41	<i>Medicago truncatula</i>
35	G663	7673087	5.10E-43	<i>Petunia integrifolia</i>
35	G663	9508051	3.00E-41	<i>Lycopersicon esculentum</i>
35	G663	7673091	3.30E-41	<i>Petunia x hybrida</i>
35	G663	7673097	2.40E-36	<i>Petunia axillaris</i>
35	G663	5048991	1.20E-33	<i>Gossypium hirsutum</i>
35	G663	6455590	2.50E-31	<i>Glycine max</i>
35	G663	7560175	1.90E-27	<i>Medicago truncatula</i>
35	G663	7244424	4.10E-26	<i>Mentha x piperita</i>
35	G663	9954117	3.40E-25	<i>Solanum tuberosum</i>
35	G663	6020191	3.60E-25	<i>Pinus taeda</i>
37	G1396	498704	5.20E-22	<i>Spinacia oleracea</i>
37	G1396	7502400	1.20E-21	<i>Gossypium arboreum</i>
37	G1396	3857536	3.40E-21	<i>Populus balsamifera subsp. trichocarpa</i>
37	G1396	4385300	1.20E-20	<i>Lycopersicon esculentum</i>
37	G1396	6917249	1.50E-20	<i>Lycopersicon pennellii</i>
37	G1396	6915979	1.70E-20	<i>Glycine max</i>
37	G1396	7674530	2.70E-20	<i>Medicago truncatula</i>
37	G1396	8090319	3.40E-20	<i>Sorghum bicolor</i>
37	G1396	3592182	9.10E-20	<i>Oryza sativa</i>
37	G1396	6654124	1.10E-19	<i>Zea mays</i>
39	G265	5821137	6.50E-83	<i>Nicotiana tabacum</i>
39	G265	7158881	3.80E-79	<i>Medicago sativa</i>
39	G265	886741	1.60E-70	<i>Zea mays</i>
39	G265	5900449	5.60E-43	<i>Lycopersicon esculentum</i>
39	G265	8903922	8.20E-43	<i>Hordeum vulgare</i>
39	G265	7561318	2.10E-41	<i>Medicago truncatula</i>
39	G265	9204445	5.30E-36	<i>Glycine max</i>
39	G265	4528238	5.40E-36	<i>Citrus unshiu</i>
39	G265	19489	2.10E-35	<i>Lycopersicon peruvianum</i>
39	G265	9251913	2.00E-32	<i>Solanum tuberosum</i>
41	G1006	7528275	2.70E-51	<i>Mesembryanthemum crystallinum</i>
41	G1006	3342210	4.90E-49	<i>Lycopersicon esculentum</i>
41	G1006	6654776	1.90E-48	<i>Medicago truncatula</i>
41	G1006	790362	2.30E-47	<i>Nicotiana tabacum</i>
41	G1006	8809570	2.00E-46	<i>Nicotiana sylvestris</i>
41	G1006	7627061	6.40E-41	<i>Gossypium arboreum</i>
41	G1006	7324479	1.20E-35	<i>Lycopersicon pennellii</i>
41	G1006	6478844	1.80E-35	<i>Matricaria chamomilla</i>

Figure 3E

SEQ ID No.	GID	Genbank NID	P-value	Species
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43	G1309	7566043	9.60E-35	Medicago truncatula
43	G1309	5891104	2.20E-31	Lycopersicon esculentum
43	G1309	5860031	2.10E-30	Pinus taeda
43	G1309	5049507	6.20E-30	Gossypium hirsutum
43	G1309	5139805	1.30E-29	Glycine max
43	G1309	6850206	2.50E-29	Oryza sativa
43	G1309	7721017	3.40E-29	Lotus japonicus
43	G1309	8368245	5.20E-28	Zea mays
43	G1309	20560	9.50E-27	Petunia x hybrida
45	G2550	4380729	2.80E-51	Lycopersicon esculentum
45	G2550	5667196	2.20E-49	Oryza sativa
45	G2550	8669454	1.40E-48	Glycine max
45	G2550	9298824	1.50E-48	Sorghum bicolor
45	G2550	7239156	9.90E-46	Malus x domestica
45	G2550	7570704	5.70E-45	Medicago truncatula
45	G2550	7628137	3.30E-42	Gossypium arboreum
45	G2550	7244366	6.00E-41	Mentha x piperita
45	G2550	9428023	4.70E-40	Triticum aestivum
45	G2550	9250642	3.50E-39	Solanum tuberosum
47	G965	7239156	3.10E-126	Malus x domestica
47	G965	5892190	2.00E-62	Lycopersicon esculentum
47	G965	7628137	1.60E-56	Gossypium arboreum
47	G965	9205496	2.60E-49	Glycine max
47	G965	6069643	1.70E-45	Oryza sativa
47	G965	7562931	2.50E-44	Medicago truncatula
47	G965	7781695	1.60E-41	Lotus japonicus
47	G965	9298824	6.30E-33	Sorghum bicolor
47	G965	9428023	1.50E-31	Triticum aestivum
47	G965	7244366	1.20E-29	Mentha x piperita
49	G793	6976712	3.60E-43	Lycopersicon esculentum
49	G793	7924288	2.00E-41	Glycine max
49	G793	7614163	3.90E-34	Lotus japonicus
49	G793	9198126	5.70E-23	Medicago truncatula
49	G793	5804781	1.10E-22	Nicotiana tabacum
49	G793	7535578	1.60E-14	Sorghum bicolor
49	G793	4276777	6.10E-14	Oryza sativa
49	G793	5915205	2.90E-10	Zea mays
49	G793	9249806	4.20E-10	Solanum tuberosum
49	G793	7624745	1.30E-09	Gossypium arboreum
51	G764	4384535	7.00E-70	Lycopersicon esculentum
51	G764	5049217	1.80E-65	Gossypium hirsutum
51	G764	6454868	1.90E-64	Glycine max
51	G764	6066594	5.20E-59	Petunia x hybrida
51	G764	4218536	2.30E-52	Triticum sp.
51	G764	6732159	2.30E-52	Triticum monococcum
51	G764	9361647	7.50E-52	Triticum aestivum
51	G764	4977542	4.10E-49	Oryza sativa
51	G764	6799764	4.40E-49	Medicago truncatula
51	G764	9296257	1.00E-48	Sorghum bicolor

Figure 3F

SEQ ID No.	GID	Genbank NID	P-value	Species
53	G350	439492	5.20E-53	<i>Petunia x hybrida</i>
53	G350	7228328	8.90E-51	<i>Medicago sativa</i>
53	G350	4666359	3.10E-48	<i>Datisca glomerata</i>
53	G350	1763062	8.30E-48	<i>Glycine max</i>
53	G350	7626808	9.10E-44	<i>Gossypium arboreum</i>
53	G350	7206360	2.20E-43	<i>Medicago truncatula</i>
53	G350	2981168	2.10E-38	<i>Nicotiana tabacum</i>
53	G350	7322653	2.00E-37	<i>Lycopersicon hirsutum</i>
53	G350	5276755	2.40E-37	<i>Lycopersicon esculentum</i>
53	G350	2058503	1.10E-31	<i>Brassica rapa</i>
55	G986	6472584	1.00E-34	<i>Nicotiana tabacum</i>
55	G986	8440065	8.80E-33	<i>Gossypium hirsutum</i>
55	G986	4385167	1.50E-32	<i>Lycopersicon esculentum</i>
55	G986	8205146	5.50E-30	<i>Oryza sativa</i>
55	G986	5820418	8.80E-26	<i>Glycine max</i>
55	G986	1159878	2.30E-23	<i>Avena fatua</i>
55	G986	9250698	4.60E-22	<i>Solanum tuberosum</i>
55	G986	9413507	7.90E-21	<i>Triticum aestivum</i>
55	G986	7748539	2.30E-20	<i>Lotus japonicus</i>
55	G986	9199620	1.30E-16	<i>Medicago truncatula</i>
57	G1349	8904043	1.50E-47	<i>Hordeum vulgare</i>
57	G1349	7244408	2.40E-47	<i>Mentha x piperita</i>
57	G1349	8286031	3.60E-46	<i>Glycine max</i>
57	G1349	9298238	9.10E-36	<i>Sorghum bicolor</i>
57	G1349	7628908	4.70E-34	<i>Gossypium arboreum</i>
57	G1349	5046180	1.50E-33	<i>Gossypium hirsutum</i>
57	G1349	5888938	1.30E-30	<i>Lycopersicon esculentum</i>
57	G1349	5043924	6.20E-30	<i>Pinus taeda</i>
57	G1349	8845076	4.40E-29	<i>Triticum aestivum</i>
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## MBI15 Sequence Listing.ST25

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aat tct cat cct ttc tca atc tcc gat cat cat cat cat cat cct Asn Ser His Pro Phe Ser Ser Ile Ser Asp His His His His His Pro 225 230 235	842
cat cat cag cat caa gag ttt tca ttc gtt ccc gac cat ttg ata tca His His Gln His Glu Phe Ser Phe Val Pro Asp His Leu Ile Ser 240 245 250	890
ccg gca gaa tcc aac ggc gga gca ttc aat ctt gat ttt aat atg tca Pro Ala Glu Ser Asn Gly Gly Ala Phe Asn Leu Asp Phe Asn Met Ser 255 260 265 270	938
aca ccc tcc ggc gcc gga gct gcc gtc tcc gcc gca tca ggt ggt ggc Thr Pro Ser Gly Ala Gly Ala Val Ser Ala Ala Ser Gly Gly Gly 275 280 285	986
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gga gga ggt cca cag ttc tta ttc ggt gca ctg cct gca gag aat cac Gly Gly Pro Gln Phe Leu Phe Gly Ala Leu Pro Ala Glu Asn His 320 325 330	1130
cac cac aat cac cag ttt cag ctt tac tat gaa aat gga tgc aga aac His His Asn His Gln Phe Gln Leu Tyr Tyr Glu Asn Gly Cys Arg Asn 335 340 345 350	1178
tca tca gaa cat aag ggt aaa ggc aag aac tga tgatattaat tattgcattct Ser Ser Glu His Lys Gly Lys Gly Lys Asn 355 360	1231
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Gly Pro Arg Asp Arg Arg Val Arg Leu Ser Val Ser Thr Ala Leu Gln 50 55 60	
Phe Tyr Asp Leu Gln Asp Arg Leu Gly Tyr Asp Gln Pro Ser Lys Ala 65 70 75 80	

## MBI15 Sequence Listing.ST25

Val Glu Trp Leu Ile Lys Ala Ala Glu Asp Ser Ile Ser Glu Leu Pro  
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Ser Leu Asn Asn Thr His Phe Pro Thr Asp Asp Glu Asn His Gln Asn  
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Gln Thr Leu Thr Thr Val Ala Ala Asn Ser Leu Ser Lys Ser Ala Cys  
115 120 125

Ser Ser Asn Ser Asp Thr Ser Lys Asn Ser Ser Gly Leu Ser Leu Ser  
130 135 140

Arg Ser Glu Leu Arg Asp Lys Ala Arg Glu Arg Ala Arg Glu Arg Thr  
145 150 155 160

Ala Lys Glu Thr Lys Glu Arg Asp His Asn His Thr Ser Phe Thr Asp  
165 170 175

Leu Leu Asn Ser Gly Ser Asp Pro Val Asn Ser Asn Arg Gln Trp Met  
180 185 190

Ala Ser Ala Pro Ser Ser Ser Pro Met Glu Tyr Phe Ser Ser Gly Leu  
195 200 205

Ile Leu Gly Ser Gly Gln Gln Thr His Phe Pro Ile Ser Thr Asn Ser  
210 215 220

His Pro Phe Ser Ser Ile Ser Asp His His His His His Pro His His  
225 230 235 240

Gln His Gln Glu Phe Ser Phe Val Pro Asp His Leu Ile Ser Pro Ala  
245 250 255

Glu Ser Asn Gly Gly Ala Phe Asn Leu Asp Phe Asn Met Ser Thr Pro  
260 265 270

Ser Gly Ala Gly Ala Ala Val Ser Ala Ala Ser Gly Gly Phe Ser  
275 280 285

Gly Phe Asn Arg Gly Thr Leu Gln Ser Asn Ser Thr Asn Gln His Gln  
290 295 300

Ser Phe Leu Ala Asn Leu Gln Arg Phe Pro Thr Ser Glu Ser Gly Gly  
305 310 315 320

Gly Pro Gln Phe Leu Phe Gly Ala Leu Pro Ala Glu Asn His His  
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## MBI15 Sequence Listing .ST25

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## MBI15 Sequence Listing.ST25

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His Pro Thr Asn Gln Val Asn Val Lys Glu Glu Ala Val Lys Lys Glu		
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Gln Ala Thr Glu Pro Gly Lys Arg Arg Lys Arg Lys Asn Val Tyr Arg		
65	70	75
Gly Ile Arg Lys Arg Pro Trp Gly Lys Trp Ala Ala Glu Ile Arg Asp		
85	90	95
Pro Arg Lys Gly Val Arg Val Trp Leu Gly Thr Phe Asn Thr Ala Glu		
100	105	110
Glu Ala Ala Met Ala Tyr Asp Val Ala Ala Lys Gln Ile Arg Gly Asp		
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Lys Ala Lys Leu Asn Phe Pro Asp Leu His His Pro Pro Pro Asn		
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Tyr Thr Pro Pro Pro Ser Ser Pro Arg Ser Thr Asp Gln Pro Pro Ala		
145	150	155
160		
Lys Lys Val Cys Val Val Ser Gln Ser Glu Ser Glu Leu Ser Gln Pro		
165	170	175
Ser Phe Pro Val Glu Cys Ile Gly Phe Gly Asn Gly Asp Glu Phe Gln		
180	185	190
Asn Leu Ser Tyr Gly Phe Glu Pro Asp Tyr Asp Leu Lys Gln Gln Ile		
195	200	205
Ser Ser Leu Glu Ser Phe Leu Glu Leu Asp Gly Asn Thr Ala Glu Gln		
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Pro Ser Gln Leu Asp Glu Ser Val Ser Glu Val Asp Met Trp Met Leu		

MBI15 Sequence Listing.ST25

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gcttttgc	tattgtctt	tat tagaaa	cagtggtag	tttttagtct	ttcacattgt	180		
tcaagttcga	agctttttt	ggagggaatt	ttgggcttct	gatttgatc	gaaacttact	240		
gatagtaagt	tcttgagtc	ctcccttaact	gtatgttctg	tgtactgaag	ttattgaatt	300		
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			Met	Asp	Glu	Asn	Asn	His
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Met Val Asp Asp Ser Ser Asp Ser Ile Val Ser Trp Ser Gln Ser								
25	30	35						
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Asn Lys Ser Phe Ile Val Trp Asn Pro Pro Glu Phe Ser Arg Asp Leu								
40	45	50						
ctt ccg aga ttc ttc aag cac aat aac ttc tct agc ttt atc cgc cag						667		
Leu Pro Arg Phe Phe Lys His Asn Asn Phe Ser Ser Phe Ile Arg Gln								
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ctt aac aca tat ggt ttt aga aaa gct gat cct gag caa tgg gaa ttt						715		
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75	80	85						
gcg aat gat gat ttt gtg aga ggt caa cct cat ctt atg aag aac att						763		
Ala Asn Asp Asp Phe Val Arg Gly Gln Pro His Leu Met Lys Asn Ile								
90	95	100						
cat aga cgc aaa cca gtt cat agc cac tct tta ccg aat ctt caa gct						811		
His Arg Arg Lys Pro Val His Ser His Ser Leu Pro Asn Leu Gln Ala								
105	110	115						
cag tta aac ccg ttg acg gat tca gaa cga gtg aga atg aat aat cag						859		
Gln Leu Asn Pro Leu Thr Asp Ser Glu Arg Val Arg Met Asn Asn Gln								
120	125	130						
att gag aga ttg aca aaa gag aaa gaa gga ttg ctt gaa gag tta cat						907		
Ile Glu Arg Leu Thr Lys Glu Lys Glu Gly Leu Leu Glu Glu Leu His								
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aaa caa gac gag gaa cga gaa gtg ttt gag atg caa gtg aaa gaa ctt						955		
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MBI15 Sequence Listing.ST25

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ccg tgt gtt ccc gaa aca aac gag agg aaa aga agg ttc cct agg atc Pro Cys Val Pro Glu Thr Asn Glu Arg Lys Arg Arg Phe Pro Arg Ile	200	205	1099
gag ttc ttt ccc gat gaa ccg atg ttg gaa gag aac aaa act tgt gtt Glu Phe Pro Asp Glu Pro Met Leu Glu Glu Asn Lys Thr Cys Val	215	220	1147
gtt gtg aga gag gaa ggt tct aca agc cct tct tca cac aca aga gag Val Val Arg Glu Glu Gly Ser Thr Ser Pro Ser Ser His Thr Arg Glu	235	240	1195
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gta tcg gat tct tgt gag agt atg tta caa tca aga agt atg atg aca Val Ser Asp Ser Cys Glu Ser Met Leu Gln Ser Arg Ser Met Met Thr	265	270	1291
ctt gat gtg gat gaa tca tct act ttt cca gag agc cct cct ctt tct Leu Asp Val Asp Glu Ser Ser Thr Phe Pro Glu Ser Pro Pro Leu Ser	280	285	1339
tgc ata cag tta agt gtc gat tca cgt ctc aaa tct cct cct tct cca Cys Ile Gln Leu Ser Val Asp Ser Arg Leu Lys Ser Pro Pro Ser Pro	295	300	1387
agg atc atc gat atg aac tgt gag ccc gat ggt tcg aaa gaa cag aac Arg Ile Ile Asp Met Asn Cys Glu Pro Asp Gly Ser Lys Glu Gln Asn	315	320	1435
act gtt gct gct cct cct cca gta gca gga gcg aat gat ggc Thr Val Ala Ala Pro Pro Pro Pro Val Ala Gly Ala Asn Asp Gly	330	335	1483
ttc tgg cag cag ttt ttc tca gag aat cct ggc tca acc gag caa cgg Phe Trp Gln Gln Phe Phe Ser Glu Asn Pro Gly Ser Thr Glu Gln Arg	345	350	1531
gaa gtt caa tta gag agg aaa gac gat aaa gat aaa gcc gga gta cgt Glu Val Gln Leu Glu Arg Lys Asp Asp Lys Asp Lys Ala Gly Val Arg	360	365	1579
act gag aaa tgt tgg tgg aat tcg aga aat gtt aat gca att aca gaa Thr Glu Lys Cys Trp Trp Asn Ser Arg Asn Val Asn Ala Ile Thr Glu	375	380	1627
cag ctt gga cat ctg act tct tca gag aga agt tga tatgtcaaag Gln Leu Gly His Leu Thr Ser Ser Glu Arg Ser	395	400	1673
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## MBI15 Sequence Listing.ST25

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35 40 45

Glu Phe Ser Arg Asp Leu Leu Pro Arg Phe Phe Lys His Asn Asn Phe  
50 55 60

Ser Ser Phe Ile Arg Gln Leu Asn Thr Tyr Gly Phe Arg Lys Ala Asp  
65 70 75 80

Pro Glu Gln Trp Glu Phe Ala Asn Asp Asp Phe Val Arg Gly Gln Pro  
85 90 95

His Leu Met Lys Asn Ile His Arg Arg Lys Pro Val His Ser His Ser  
100 105 110

Leu Pro Asn Leu Gln Ala Gln Leu Asn Pro Leu Thr Asp Ser Glu Arg  
115 120 125

Val Arg Met Asn Asn Gln Ile Glu Arg Leu Thr Lys Glu Lys Glu Gly  
130 135 140

Leu Leu Glu Glu Leu His Lys Gln Asp Glu Glu Arg Glu Val Phe Glu  
145 150 155 160

Met Gln Val Lys Glu Leu Lys Glu Arg Leu Gln His Met Glu Lys Arg  
165 170 175

Gln Lys Thr Met Val Ser Phe Val Ser Gln Val Leu Glu Lys Pro Gly  
180 185 190

Leu Ala Leu Asn Leu Ser Pro Cys Val Pro Glu Thr Asn Glu Arg Lys  
195 200 205

Arg Arg Phe Pro Arg Ile Glu Phe Phe Pro Asp Glu Pro Met Leu Glu  
210 215 220

Glu Asn Lys Thr Cys Val Val Val Arg Glu Glu Gly Ser Thr Ser Pro  
225 230 235 240

Ser Ser His Thr Arg Glu His Gln Val Glu Gln Leu Glu Ser Ser Ile  
245 250 255

Ala Ile Trp Glu Asn Leu Val Ser Asp Ser Cys Glu Ser Met Leu Gln  
260 265 270

Ser Arg Ser Met Met Thr Leu Asp Val Asp Glu Ser Ser Thr Phe Pro  
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MBI15 Sequence Listing .ST25

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305															320
310                   315															

Gly	Ser	Lys	Glu	Gln	Asn	Thr	Val	Ala	Ala	Pro	Pro	Pro	Pro	Val	
325														335	
330															

Ala	Gly	Ala	Asn	Asp	Gly	Phe	Trp	Gln	Gln	Phe	Phe	Ser	Glu	Asn	Pro
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Asp	Lys	Ala	Gly	Val	Arg	Thr	Glu	Lys	Cys	Trp	Trp	Asn	Ser	Arg	Asn
370															380
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	Met	Ser	Met	Thr	Ala	Asp	Ser	Gln	Ser	Asp	Tyr	Ala	Phe	Leu	Glu
1															15

tcc	ata	cga	cga	cac	tta	cta	gga	gaa	tcg	qag	ccg	ata	ctc	agt	gag
	Ser	Ile	Arg	Arg	His	Leu	Leu	Gly	Ser	Glu	Pro	Ile	Leu	Ser	Glu
20															30

tcg	aca	gca	agt	tcg	gtt	act	caa	tct	tgt	gta	acc	ggt	cag	agc	att
	Ser	Thr	Ala	Ser	Ser	Val	Thr	Gln	Ser	Cys	Val	Thr	Gly	Gln	Ser
35															45

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Lys	Pro	Val	Tyr	Gly	Arg	Asn	Pro	Ser	Phe	Ser	Lys	Leu	Tyr	Pro	Cys
50															60

ttc	acc	gag	agc	tgg	gga	gat	ttg	ccg	ttg	aaa	gaa	aac	gat	tct	gag
Phe	Thr	Glu	Ser	Trp	Gly	Asp	Leu	Pro	Leu	Lys	Glu	Asn	Ser	Glu	
65															75

gat	atg	tta	gtt	tac	ggt	atc	ctc	aac	gac	gcc	ttt	cac	ggc	ggt	tgg
Asp	Met	Leu	Val	Tyr	Gly	Ile	Leu	Asn	Asp	Ala	Phe	His	Gly	Gly	Trp
80															95

gag	ccg	tct	tct	tcg	tct	tcc	gac	gaa	gat	cgt	agc	tct	ttc	ccg	agt
Glu	Pro	Ser	Ser	Ser	Ser	Ser	Asp	Glu	Asp	Arg	Ser	Ser	Phe	Pro	Ser
100															110

gtt	aag	atc	gag	act	ccg	gag	agt	ttc	gcg	gct	gat	tct	gtt	ccg	
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MBI15 Sequence Listing.ST25

Val Lys Ile Glu Thr Pro Glu Ser Phe Ala Ala Val Asp Ser Val Pro			
115	120	125	
gtc aag aag gag aag acg agt cct gtt tcg gcg gcg gtg acg gcg gcg		491	
Val Lys Lys Glu Lys Thr Ser Pro Val Ser Ala Ala Val Thr Ala Ala			
130	135	140	
aag gga aag cat tat aga gga gtg aga caa agg ccg tgg ggg aaa ttt		539	
Lys Gly Lys His Tyr Arg Gly Val Arg Gln Arg Pro Trp Gly Lys Phe			
145	150	155	
gcg gcg gag att aga gat ccg gcg aag aac gga gct agg gtt tgg tta		587	
Ala Ala Glu Ile Arg Asp Pro Ala Lys Asn Gly Ala Arg Val Trp Leu			
160	165	170	175
gga acg ttt gag acg gcg gag gac gcg gcg ttg gct tac gac aga gct		635	
Gly Thr Phe Glu Thr Ala Glu Asp Ala Ala Leu Ala Tyr Asp Arg Ala			
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Ala Phe Arg Met Arg Gly Ser Arg Ala Leu Leu Asn Phe Pro Leu Arg			
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Val Asn Ser Gly Glu Pro Asp Pro Val Arg Ile Lys Ser Lys Arg Ser			
210	215	220	
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Ser Phe Ser Ser Asn Glu Asn Gly Ala Pro Lys Lys Arg Arg Thr			
225	230	235	
gtg gcc gcc ggt ggt gga atg gat aag gga ttg acg gtg aag tgc gag		827	
Val Ala Ala Gly Gly Met Asp Lys Gly Leu Thr Val Lys Cys Glu			
240	245	250	255
gtt gtt gaa gtg gca cgt ggc gat cgt tta ttg gtt tta taa		869	
Val Val Glu Val Ala Arg Gly Asp Arg Leu Leu Val Leu			
260	265		
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Thr Ala Ser Ser Val Thr Gln Ser Cys Val Thr Gly Gln Ser Ile Lys			
35	40	45	
Pro Val Tyr Gly Arg Asn Pro Ser Phe Ser Lys Leu Tyr Pro Cys Phe			
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Thr Glu Ser Trp Gly Asp Leu Pro Leu Lys Glu Asn Asp Ser Glu Asp			
65	70	75	80
Met Leu Val Tyr Gly Ile Leu Asn Asp Ala Phe His Gly Gly Trp Glu			
85	90	95	

MBI15 Sequence Listing.ST25

Pro	Ser	Ser	Ser	Ser	Ser	Asp	Glu	Asp	Arg	Ser	Ser	Phe	Pro	Ser	Val
100						105						110			
Lys Ile Glu Thr Pro Glu Ser Phe Ala Ala Val Asp Ser Val Pro Val															
115						120						125			
Lys Lys Glu Lys Thr Ser Pro Val Ser Ala Ala Val Thr Ala Ala Lys															
130						135						140			
Gly Lys His Tyr Arg Gly Val Arg Gln Arg Pro Trp Gly Lys Phe Ala															
145						150						155			160
Ala Glu Ile Arg Asp Pro Ala Lys Asn Gly Ala Arg Val Trp Leu Gly															
165						170						175			
Thr Phe Glu Thr Ala Glu Asp Ala Ala Leu Ala Tyr Asp Arg Ala Ala															
180						185						190			
Phe Arg Met Arg Gly Ser Arg Ala Leu Leu Asn Phe Pro Leu Arg Val															
195						200						205			
Asn Ser Gly Glu Pro Asp Pro Val Arg Ile Lys Ser Lys Arg Ser Ser															
210						215						220			
Phe Ser Ser Ser Asn Glu Asn Gly Ala Pro Lys Lys Arg Arg Thr Val															
225						230						235			240
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Val Glu Val Ala Arg Gly Asp Arg Leu Leu Val Leu															
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CTCCGATTc ATCATCATCT TCCCCATCAT CGTCGTCTT GAAATCTGT CTTCtCAACG 180															
CTCTTCACTT CTGCTGTAAT AAGCAGAGGC TTGTTCTGGA GACTCCtCT CTTCCATGC 240															
GCTTAAGACC CAAAAGGACT TGTTCAGTG TTGAAGTCTT TGGGGGTTT CACATAAAGC 300															
AGCAAAAGTT TTCTTTTTC ATAGTTCGCT GAGAGTTTG AGTTTGATA CAAAAAAAGT 360															
TTTGACCTTT TAGAGTGATT TTTGTTCTT TCTGTTCTT GGtATTTT GAGGAGTGGG 420															
TTTAACA ATG GTT GCG ATT AGA AAG GAA CAG TCT TTG AGT GGT GTT AGT 469															
Met Val Ala Ile Arg Lys Glu Gln Ser Leu Ser Gly Val Ser															
1 5 10															
AGC GAG ATT AAG AAG AGA GCT AAG AGA AAC ACT CTA TCG TCC CTT CCT 517															

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Ser	Glu	Ile	Lys	Lys	Arg	Ala	Lys	Arg	Asn	Thr	Leu	Ser	Ser	Leu	Pro
15					20					25					30
caa gaa acc caa cct ttg agg aaa gtc cgt att att gtg aat gat cct															565
Gln Glu Thr Gln Pro Leu Arg Lys Val Arg Ile Ile Val Asn Asp Pro															
35 40 45															
tat gct act gat gat tcc tct agt gat gag gaa gag ctt aag gtt cct															613
Tyr Ala Thr Asp Asp Ser Ser Asp Glu Glu Glu Leu Lys Val Pro															
50 55 60															
aag cca agg aaa atg aaa cgt atc gtt cgt gag att aac ttt cct tct															661
Lys Pro Arg Lys Met Lys Arg Ile Val Arg Glu Ile Asn Phe Pro Ser															
65 70 75															
atg gaa gtt tct gaa cag cct tct gag agt tct tct cag gac agt act															709
Met Glu Val Ser Glu Gln Pro Ser Glu Ser Ser Gln Asp Ser Thr															
80 85 90															
aaa act gat ggc aag ata gct gtg tca gct tct cct gct gtt cct agg															757
Lys Thr Asp Gly Lys Ile Ala Val Ser Ala Ser Pro Ala Val Pro Arg															
95 100 105 110															
aag aag cct gtt ggt gtt agg caa agg aaa tgg ggg aaa tgg gct gct															805
Lys Lys Pro Val Gly Val Arg Gln Arg Lys Trp Gly Lys Trp Ala Ala															
115 120 125															
gag att aga gat cct att aag aaa act agg act tgg ttg ggt act ttt															853
Glu Ile Arg Asp Pro Ile Lys Lys Thr Arg Thr Trp Leu Gly Thr Phe															
130 135 140															
gat act ctt gaa gaa gct gct aaa gct tat gat gct aag aag ctt gag															901
Asp Thr Leu Glu Ala Ala Lys Ala Tyr Asp Ala Lys Lys Leu Glu															
145 150 155															
ttt gat gct att gtt gct gga aat gtg tcc act act aaa cgt gat gtt															949
Phe Asp Ala Ile Val Ala Gly Asn Val Ser Thr Thr Lys Arg Asp Val															
160 165 170															
tct tca tct gag act agc caa tgc tct cgt tct tca cct gtt gtt cct															997
Ser Ser Ser Glu Thr Ser Gln Cys Ser Arg Ser Pro Val Val Pro															
175 180 185 190															
gtt gag caa gat gac act tct gca tca gct ctc act tgg gtc aac aac															1045
Val Glu Gln Asp Asp Thr Ser Ala Ser Ala Leu Thr Cys Val Asn Asn															
195 200 205															
cct gat gac gtc tcg acc gtt gct cca act gct cca act cca aat gtt															1093
Pro Asp Asp Val Ser Thr Val Ala Pro Thr Ala Pro Thr Pro Asn Val															
210 215 220															
cct gct ggt gga aac aag gaa acg ttg ttc gat ttc gac ttt act aat															1141
Pro Ala Gly Gly Asn Lys Glu Thr Leu Phe Asp Phe Asp Phe Thr Asn															
225 230 235															
cta cag atc cct gat ttt ggt ttg ttg gca gag gag caa caa gac cta															1189
Leu Gln Ile Pro Asp Phe Gly Phe Leu Ala Glu Glu Gln Gln Asp Leu															
240 245 250															
gac ttg gat tgt ttg ctc gcg gat gat cag ttt gat gat ttg ggc ttg															1237
Asp Phe Asp Cys Phe Leu Ala Asp Asp Gln Phe Asp Asp Phe Gly Leu															
255 260 265 270															
ctt gat gac att caa gga ttc gaa gat aac ggt cca agt gcg tta cca															1285
Leu Asp Asp Ile Gln Gly Phe Glu Asp Asn Gly Pro Ser Ala Leu Pro															
275 280 285															
gat ttg gac ttt gcg gat gtt gaa gat ctt cag cta gct gac tct agt															1333
Asp Phe Asp Phe Ala Asp Val Glu Asp Leu Gln Leu Ala Asp Ser Ser															
290 295 300															
ttc ggt ttg ctt gat caa ctt gct cct atc aac atc tct tgc cca tta															1381
Phe Gly Phe Leu Asp Gln Leu Ala Pro Ile Asn Ile Ser Cys Pro Leu															
305 310 315															

MBI15 Sequence Listing.ST25

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Lys Ser Phe Ala Ala Ser	
320	
gtgtttgtt tttcgaaaa tgcttttagta atttaagaca tacaaaagtg tgtgttccgg	1492
attgttagtaa gatcttaaga cataaaagccg ggttttgcaa ttaggaatcg agtttaatg	1552
aagttttagt ttatgtttg	1571
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Thr Gln Pro Leu Arg Lys Val Arg Ile Ile Val Asn Asp Pro Tyr Ala	
35 40 45	
Thr Asp Asp Ser Ser Ser Asp Glu Glu Leu Lys Val Pro Lys Pro	
50 55 60	
Arg Lys Met Lys Arg Ile Val Arg Glu Ile Asn Phe Pro Ser Met Glu	
65 70 75 80	
Val Ser Glu Gln Pro Ser Glu Ser Ser Gln Asp Ser Thr Lys Thr	
85 90 95	
Asp Gly Lys Ile Ala Val Ser Ala Ser Pro Ala Val Pro Arg Lys Lys	
100 105 110	
Pro Val Gly Val Arg Gln Arg Lys Trp Gly Lys Trp Ala Ala Glu Ile	
115 120 125	
Arg Asp Pro Ile Lys Lys Thr Arg Thr Trp Leu Gly Thr Phe Asp Thr	
130 135 140	
Leu Glu Glu Ala Ala Lys Ala Tyr Asp Ala Lys Lys Leu Glu Phe Asp	
145 150 155 160	
Ala Ile Val Ala Gly Asn Val Ser Thr Thr Lys Arg Asp Val Ser Ser	
165 170 175	
Ser Glu Thr Ser Gln Cys Ser Arg Ser Ser Pro Val Val Pro Val Glu	
180 185 190	
Gln Asp Asp Thr Ser Ala Ser Ala Leu Thr Cys Val Asn Asn Pro Asp	
195 200 205	
Asp Val Ser Thr Val Ala Pro Thr Ala Pro Thr Pro Asn Val Pro Ala	
210 215 220	
Gly Gly Asn Lys Glu Thr Leu Phe Asp Phe Asp Phe Thr Asn Leu Gln	

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225	230	235	240													
Ile Pro Asp Phe Gly Phe Leu Ala Glu Glu Gln Gln Asp Leu Asp Phe																
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Asp Cys Phe Leu Ala Asp Asp Gln Phe Asp Asp Phe Gly Leu Leu Asp																
260		265	270													
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275		280	285													
Asp Phe Ala Asp Val Glu Asp Leu Gln Leu Ala Asp Ser Ser Phe Gly																
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1		5			10				15							
cct	gaa	gaa	gac	gag	aag	cta	agg	agc	ttc	atc	ctc	tct	tat	ggc	cat	96
Pro	Glu	Glu	Asp	Glu	Lys	Leu	Arg	Ser	Phe	Ile	Leu	Ser	Tyr	Gly	His	
20		25		25		30										
tct	tgc	tgg	acc	act	gtt	ccc	atc	aaa	gct	ggg	tta	caa	agg	aat	ggg	144
Ser	Cys	Trp	Thr	Thr	Val	Pro	Ile	Lys	Ala	Gly	Leu	Gln	Arg	Asn	Gly	
35		40		40		45										
aag	agc	tgc	aga	tta	aga	tgg	att	aat	tac	cta	aga	cca	ggg	tta	aag	192
Lys	Ser	Cys	Arg	Leu	Arg	Trp	Ile	Asn	Tyr	Leu	Arg	Pro	Gly	Leu	Lys	
50		55		55		60										
agg	gat	att	agt	gca	gaa	gaa	gag	act	atc	ttg	acg	ttt	cat		240	
Arg	Asp	Met	Ile	Ser	Ala	Glu	Glu	Glu	Thr	Ile	Leu	Thr	Phe	His		
65		70		70		75		75		80						
tct	ccc	ttg	ggt	aac	aag	tgg	tgc	caa	ata	gct	aaa	ttc	tta	ccg	gga	288
Ser	Pro	Leu	Gly	Asn	Lys	Trp	Ser	Gln	Ile	Ala	Lys	Phe	Leu	Pro	Gly	
85		90		90		95										
aga	aca	gac	aat	gag	ata	aag	aac	tat	tgg	cac	tct	cat	ttg	aaa	aag	336
Arg	Thr	Asp	Asn	Glu	Ile	Lys	Asn	Tyr	Trp	His	Ser	His	Leu	Lys	Lys	
100		105		105		110										
aaa	tgg	ctc	aag	tct	cag	agc	tta	caa	gat	gca	aaa	tct	att	tcc	cct	384
Lys	Trp	Leu	Lys	Ser	Gln	Ser	Leu	Gln	Asp	Ala	Lys	Ser	Ile	Ser	Pro	
115		120		120		125										
cct	tgc	tct	tca	tca	tca	tca	ctt	gtt	gct	tgt	gga	gaa	aga	aat	ccg	432
Pro	Ser	Ser	Ser	Ser	Ser	Ser	Leu	Val	Ala	Cys	Gly	Glu	Arg	Asn	Pro	
130		135		135		140										

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gaa acc ttg atc tcg aat cac gtg ttc tcc ctc cag aga ctt cta gag Glu Thr Leu Ile Ser Asn His Val Phe Ser Leu Gln Arg Leu Leu Glu 145 150 155 160	480
aac aaa tct tca tct ccc tca caa gaa agc aac gga aat aac agc cat Asn Lys Ser Ser Ser Pro Ser Gln Glu Ser Asn Gly Asn Asn Ser His 165 170 175	528
caa tgt tct tct gct cct gag att cca agg ctt ttc ttc tct gaa tgg Gln Cys Ser Ser Ala Pro Glu Ile Pro Arg Leu Phe Phe Ser Glu Trp 180 185 190	576
ctt tct tct tca tat ccc cac acc gat tat tcc tct gag ttt acc gac Leu Ser Ser Tyr Pro His Thr Asp Tyr Ser Ser Glu Phe Thr Asp 195 200 205	624
tct aag cac agt caa gct cca aat gtc gaa gag act ctc tca gct tat Ser Lys His Ser Gln Ala Pro Asn Val Glu Glu Thr Leu Ser Ala Tyr 210 215 220	672
gaa gaa atg ggt gat gtt gat cag ttc cat tac aac gaa atg atg atc Glu Glu Met Gly Asp Val Asp Gln Phe His Tyr Asn Glu Met Met Ile 225 230 235 240	720
aac aac agc aac tgg act ctt aac gac att gtg ttt ggt tcc aaa tgt Asn Asn Ser Asn Trp Thr Leu Asn Asp Ile Val Phe Gly Ser Lys Cys 245 250 255	768
aag aag cag gag cat cat att tat aga gag gct tca gat tgt aat tct Lys Lys Gln Glu His His Ile Tyr Arg Glu Ala Ser Asp Cys Asn Ser 260 265 270	816
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Pro Glu Glu Asp Glu Lys Leu Arg Ser Phe Ile Leu Ser Tyr Gly His 20 25 30	
Ser Cys Trp Thr Thr Val Pro Ile Lys Ala Gly Leu Gln Arg Asn Gly 35 40 45	
Lys Ser Cys Arg Leu Arg Trp Ile Asn Tyr Leu Arg Pro Gly Leu Lys 50 55 60	
Arg Asp Met Ile Ser Ala Glu Glu Glu Thr Ile Leu Thr Phe His 65 70 75 80	
Ser Pro Leu Gly Asn Lys Trp Ser Gln Ile Ala Lys Phe Leu Pro Gly 85 90 95	
Arg Thr Asp Asn Glu Ile Lys Asn Tyr Trp His Ser His Leu Lys Lys 100 105 110	
Lys Trp Leu Lys Ser Gln Ser Leu Gln Asp Ala Lys Ser Ile Ser Pro	

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115		120	125
 Pro Ser Ser Ser Ser Ser Leu Val Ala Cys Gly Glu Arg Asn Pro 130   135                           140			
 Glu Thr Leu Ile Ser Asn His Val Phe Ser Leu Gln Arg Leu Leu Glu 145   150                           155                           160			
 Asn Lys Ser Ser Ser Pro Ser Gln Glu Ser Asn Gly Asn Asn Ser His 165   170                           175			
 Gln Cys Ser Ser Ala Pro Glu Ile Pro Arg Leu Phe Phe Ser Glu Trp 180   185                           190			
 Leu Ser Ser Ser Tyr Pro His Thr Asp Tyr Ser Ser Glu Phe Thr Asp 195   200                           205			
 Ser Lys His Ser Gln Ala Pro Asn Val Glu Glu Thr Leu Ser Ala Tyr 210   215                           220			
 Glu Glu Met Gly Asp Val Asp Gln Phe His Tyr Asn Glu Met Met Ile 225   230                           235                           240			
 Asn Asn Ser Asn Trp Thr Leu Asn Asp Ile Val Phe Gly Ser Lys Cys 245   250                           255			
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tagttgttgt tttatctg ttgtcaaaa atg gaa tcc aat tcg ttt ttc                         354			
Met Glu Ser Asn Ser Phe Phe Phe 1   5			
 gat cca tct gct tca cac ggc aac agc atg ttc ttc ctt ggg aat ctc                 402			
Asp Pro Ser Ala Ser His Gly Asn Ser Met Phe Phe Leu Gly Asn Leu 10   15                                   20			
 aat ccc gtc gtc caa gga gga gca aga tcg atg aac atg gag                         450			
Asn Pro Val Val Gln Gly Gly Ala Arg Ser Met Met Asn Met Glu			

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25	30	35	40	
gaa act tcg aag cga agg ccc ttc ttt agc tcc cct gag gat ctc tac				498
Glu Thr Ser Lys Arg Arg Pro Phe Phe Ser Ser Pro Glu Asp Leu Tyr				
45	50	55		
gac gat gac ttt tac gac gac cag ttg cct gaa aag aag cgt cgc ctc				546
Asp Asp Asp Phe Tyr Asp Asp Gln Leu Pro Glu Lys Lys Arg Arg Leu				
60	65	70		
act acc gaa caa gtg cat ctg ctg gag aaa agc ttc gag aca gag aac				594
Thr Thr Glu Gln Val His Leu Leu Glu Lys Ser Phe Glu Thr Glu Asn				
75	80	85		
aag cta gag cct gaa cgc aag act cag ctt gcc aag aag ctt ggt cta				642
Lys Leu Glu Pro Glu Arg Lys Thr Gln Leu Ala Lys Lys Leu Gly Leu				
90	95	100		
cag cca agg caa gtg gct gtc tgg ttt cag aat cgc cga gct cgt tgg				690
Gln Pro Arg Gln Val Ala Val Trp Phe Gln Asn Arg Arg Ala Arg Trp				
105	110	115	120	
aaa aca aaa cag ctt gag aga gac tac gat ctt ctc aag tcc act tac				738
Lys Thr Lys Gln Leu Glu Arg Asp Tyr Asp Leu Leu Lys Ser Thr Tyr				
125	130	135		
gac caa ctt ctt tct aac tac gac tcc atc gtc atg gac aac gat aag				786
Asp Gln Leu Leu Ser Asn Tyr Asp Ser Ile Val Met Asp Asn Asp Lys				
140	145	150		
ctc aga tcc gag gtt act tcc ctg acc gaa aag ctt cag ggc aaa caa				834
Leu Arg Ser Glu Val Thr Ser Leu Thr Glu Lys Leu Gln Gly Lys Gln				
155	160	165		
gag aca gct aat gaa cca cct ggt caa gtg ccc gaa cca aac caa ctt				882
Glu Thr Ala Asn Glu Pro Pro Gly Gln Val Pro Glu Pro Asn Gln Leu				
170	175	180		
gat ccg gtt tac att aat gcg gca gca atc aaa acc gag gac cgg tta				930
Asp Pro Val Tyr Ile Asn Ala Ala Ile Lys Thr Glu Asp Arg Leu				
185	190	195	200	
agt tca ggg agc gtt ggg agc gcg gta cta gac gac gac gca cct caa				978
Ser Ser Gly Ser Val Gly Ser Ala Val Leu Asp Asp Asp Ala Pro Gln				
205	210	215		
cta cta gac agc tgt gac tct tac ttc cca agc atc gta ccc atc caa				1026
Leu Leu Asp Ser Cys Asp Ser Tyr Phe Pro Ser Ile Val Pro Ile Gln				
220	225	230		
gac aac agc aac gcc agt gat cat gac aat gac cgg agc tgt ttc gcc				1074
Asp Asn Ser Asn Ala Ser Asp His Asp Asn Asp Arg Ser Cys Phe Ala				
235	240	245		
gac gtc ttt gtg ccc acc act tca ccg tcg cac gat cat cac ggt gaa				1122
Asp Val Phe Val Pro Thr Thr Ser Pro Ser His Asp His His Gly Glu				
250	255	260		
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Ser Leu Ala Phe Trp Gly Trp Pro				
265	270			
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## MBI15 Sequence Listing.ST25

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 20 25 30

Ala Arg Ser Met Met Asn Met Glu Glu Thr Ser Lys Arg Arg Pro Phe  
 35 40 45

Phe Ser Ser Pro Glu Asp Leu Tyr Asp Asp Asp Phe Tyr Asp Asp Gln  
 50 55 60

Leu Pro Glu Lys Lys Arg Arg Leu Thr Thr Glu Gln Val His Leu Leu  
 65 70 75 80

Glu Lys Ser Phe Glu Thr Glu Asn Lys Leu Glu Pro Glu Arg Lys Thr  
 85 90 95

Gln Leu Ala Lys Lys Leu Gly Leu Gln Pro Arg Gln Val Ala Val Trp  
 100 105 110

Phe Gln Asn Arg Arg Ala Arg Trp Lys Thr Lys Gln Leu Glu Arg Asp  
 115 120 125

Tyr Asp Leu Leu Lys Ser Thr Tyr Asp Gln Leu Leu Ser Asn Tyr Asp  
 130 135 140

Ser Ile Val Met Asp Asn Asp Lys Leu Arg Ser Glu Val Thr Ser Leu  
 145 150 155 160

Thr Glu Lys Leu Gln Gly Lys Gln Glu Thr Ala Asn Glu Pro Pro Gly  
 165 170 175

Gln Val Pro Glu Pro Asn Gln Leu Asp Pro Val Tyr Ile Asn Ala Ala  
 180 185 190

Ala Ile Lys Thr Glu Asp Arg Leu Ser Ser Gly Ser Val Gly Ser Ala  
 195 200 205

Val Leu Asp Asp Asp Ala Pro Gln Leu Leu Asp Ser Cys Asp Ser Tyr  
 210 215 220

Phe Pro Ser Ile Val Pro Ile Gln Asp Asn Ser Asn Ala Ser Asp His  
 225 230 235 240

Asp Asn Asp Arg Ser Cys Phe Ala Asp Val Phe Val Pro Thr Thr Ser  
 245 250 255

Pro Ser His Asp His His Gly Glu Ser Leu Ala Phe Trp Gly Trp Pro  
 260 265 270

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## MBI15 Sequence Listing.ST25

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&lt;222&gt; (103)..(2322)

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1	
aaa act tct cct aat act aca att ctc ttg aag act ttt cac aat aat Lys Thr Ser Pro Asn Thr Thr Ile Leu Leu Lys Thr Phe His Asn Asn	162
5 10 15 20	
tct atg tcc caa gat tat cat cat cat cat cat aat caa cac caa Ser Met Ser Gln Asp Tyr His His His His His Asn Gln His Gln	210
25 30 35	
gga ggt atc ttc aac ttc tct aat gga ttc gac cga tca gat tct ccc Gly Gly Ile Phe Asn Phe Ser Asn Gly Phe Asp Arg Ser Asp Ser Pro	258
40 45 50	
aat tta aca act cag cag aag caa gag cat caa agg gta gag atg gac Asn Leu Thr Thr Gln Gln Lys Gln Glu His Gln Arg Val Glu Met Asp	306
55 60 65	
gag gaa tct tca gtc gcc gga ggt agg att ccg gtc tac gaa tca gcc Glu Glu Ser Ser Val Ala Gly Gly Arg Ile Pro Val Tyr Glu Ser Ala	354
70 75 80	
ggt atg tta tcc gaa atg ttt aat ttc ccc gga agc agc ggt gga gga Gly Met Leu Ser Glu Met Phe Asn Phe Pro Gly Ser Ser Gly Gly Gly	402
85 90 95 100	
aga gat ctc gac ctc ggc caa tct ttc ccg tca aat agg cag ttg ctt Arg Asp Leu Asp Leu Gly Gln Ser Phe Arg Ser Asn Arg Gln Leu Leu	450
105 110 115	
gag gag caa cat cag aat att ccg gct atg aat gct acg gat tca gcc Glu Glu Gln His Asn Ile Pro Ala Met Asn Ala Thr Asp Ser Ala	498
120 125 130	
acc gcc acc gca gcc gcc atg cag tta ttc ttg atg aat cca ccg cca Thr Ala Thr Ala Ala Met Gln Leu Phe Leu Met Asn Pro Pro Pro	546
135 140 145	
ccg caa caa cca ccg tct ccg tca tcc aca act tcc cca agg agc cac Pro Gln Gln Pro Pro Ser Pro Ser Ser Thr Thr Ser Pro Arg Ser His	594
150 155 160	
cac aat tct tca act ctt cac atg tta ctt cca agt cca tcc acc aac His Asn Ser Ser Thr Leu His Met Leu Leu Pro Ser Pro Ser Thr Asn	642
165 170 175 180	
aca act cac cat cag aac tac act aat cat atg tct atg cat cag ctt Thr Thr His His Gln Asn Tyr Thr Asn His Met Ser Met His Gln Leu	690
185 190 195	
cca cat cag cat cac caa cag ata tcg acg tgg cag tct tct ccc gat Pro His Gln His Gln Ile Ser Thr Trp Gln Ser Ser Pro Asp	738
200 205 210	
cat cat cat cat cac aac agc caa acg gag att ggg acc gtc cac His His His His Asn Ser Gln Thr Glu Ile Gly Thr Val His	786
215 220 225	
gtg gaa aac agc gga gga cac gga gga caa ggc ttg tcc tta tct ctc Val Glu Asn Ser Gly Gly His Gly Gln Gly Leu Ser Leu Ser Leu	834
230 235 240	
tca tcg tct tta gag gct gca gca aaa gcg gaa gag tat aga aac att Ser Ser Ser Leu Glu Ala Ala Lys Ala Glu Glu Tyr Arg Asn Ile	882
245 250 255 260	

## MBI15 Sequence Listing.ST25

tac tac gga gcc aat tct tct aac gca tca cct cat cat caa tac aat Tyr Tyr Gly Ala Asn Ser Ser Asn Ala Ser Pro His His Gln Tyr Asn 265 270 275	930
caa ttc aag act ctt ctt gct aat tct tct caa cat cac cat caa gta Gln Phe Lys Thr Leu Leu Ala Asn Ser Ser Gln His His Gln Val 280 285 290	978
tta aac caa ttc cga tca tct ccg gct gct tct tcc tct tcc atg gca Leu Asn Gln Phe Arg Ser Ser Pro Ala Ala Ser Ser Ser Met Ala 295 300 305	1026
gcg gtc aat atc tta aga aac tcg agg tac aca acg gcc gcg caa gag Ala Val Asn Ile Leu Arg Asn Ser Arg Tyr Thr Ala Ala Gln Glu 310 315 320	1074
ttg ttg gaa gag ttt tgt agt gtt gga aga gga ttt ttg aag aag aac Leu Leu Glu Glu Phe Cys Ser Val Gly Arg Gly Phe Leu Lys Lys Asn 325 330 335 340	1122
aaa ctt ggg aac agc tca aac cct aat act tgc ggt ggt gat ggt ggt Lys Leu Gly Asn Ser Asn Pro Asn Thr Cys Gly Gly Asp Gly Gly 345 350 355	1170
ggc agc tct cct tcg tcg gcc gga gca aac aag gag cat cct cct tta Gly Ser Ser Pro Ser Ser Ala Gly Ala Asn Lys Glu His Pro Pro Leu 360 365 370	1218
tcg gcg tct gat cgg att gag cat caa aga agg aaa gtg aaa cta ctc Ser Ala Ser Asp Arg Ile Glu His Gln Arg Arg Lys Val Lys Leu Leu 375 380 385	1266
acc atg ctt gaa gag gtg gac cga cgg tac aac cat tac tgc gag caa Thr Met Leu Glu Glu Val Asp Arg Arg Tyr Asn His Tyr Cys Glu Gln 390 395 400	1314
atg cag atg gtt gtg aac tct ttc gac ata gta atg ggc cac ggt gcg Met Gln Met Val Val Asn Ser Phe Asp Ile Val Met Gly His Gly Ala 405 410 415 420	1362
gca tta ccg tac acc gca ttg gct caa aaa gct atg tca aga cat ttt Ala Leu Pro Tyr Thr Ala Leu Ala Gln Lys Ala Met Ser Arg His Phe 425 430 435	1410
aga tgc ctt aaa gat gca gtt gcg gct cag ctt aag cag agt tgc gaa Arg Cys Leu Lys Asp Ala Val Ala Ala Gln Leu Lys Gln Ser Cys Glu 440 445 450	1458
ctt ctt ggg gac aaa gat gca gcg gga atc tct tct tcc ggg tta aca Leu Leu Gly Asp Lys Asp Ala Ala Gly Ile Ser Ser Gly Leu Thr 455 460 465	1506
aaa ggt gaa act ccg cgt ttg cgt ttg cta gag caa agt ttg cgt cag Lys Gly Glu Thr Pro Arg Leu Arg Leu Leu Glu Gln Ser Leu Arg Gln 470 475 480	1554
caa cgt gcg ttt cat caa atg ggt atg atg gaa caa gaa gct tgg cgg Gln Arg Ala Phe His Gln Met Gly Met Met Glu Gln Glu Ala Trp Arg 485 490 495 500	1602
cct caa ccg ggt ttg cct gaa cgc tcc gtc aat ata ctt aga gct tgg Pro Gln Arg Gly Leu Pro Glu Arg Ser Val Asn Ile Leu Arg Ala Trp 505 510 515	1650
ctc ttc gaa cat ttc ctt cac ccg tat cca agt gat gca gat aaa cac Leu Phe Glu His Phe Leu His Pro Tyr Pro Ser Asp Ala Asp Lys His 520 525 530	1698
cta ttg gct cga cag act ggt tta tcc aga aat cag gta tca aat tgg Leu Leu Ala Arg Gln Thr Gly Leu Ser Arg Asn Gln Val Ser Asn Trp 535 540 545	1746
ttc ata aat gct agg gtt cgt tta tgg aaa cca atg gtg gaa gaa atg Phe Ile Asn Ala Arg Val Arg Leu Trp Lys Pro Met Val Glu Glu Met	1794

MBI15 Sequence Listing .ST25

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aac gaa gaa gat caa gaa aca aaa aac agc aac gac gac aag aac aca Asn Glu Glu Asp Gln Glu Thr Lys Asn Ser Asn Asp Asp Lys Ser Thr 585 590 595			1890
aaa tcc aac aac aat gaa agc aac ttc act gcc gtt cgg acc act tca Lys Ser Asn Asn Glu Ser Asn Phe Thr Ala Val Arg Thr Thr Ser 600 605 610			1938
caa act cca acg aca acc gca cca gac gca tca gac gca gac gca Gln Thr Pro Thr Thr Ala Pro Asp Ala Ser Asp Ala Asp Ala Ala 615 620 625			1986
gta gcg aca ggc cac cgt cta aga tcc aac att aat gct tac gaa aac Val Ala Thr Gly His Arg Leu Arg Ser Asn Ile Asn Ala Tyr Glu Asn 630 635 640			2034
gac gct tca tca ctt cta ctc cct tcc tct tat tcc aac gcc gcc gct Asp Ala Ser Ser Leu Leu Pro Ser Ser Tyr Ser Asn Ala Ala Ala 645 650 655 660			2082
cct gcc gct gtt tct gac gac ttg aat tct cgt tac ggt ggc tca gac Pro Ala Ala Val Ser Asp Asp Leu Asn Ser Arg Tyr Gly Gly Ser Asp 665 670 675			2130
gcg ttt tcc gcc gtt gcc acg tgt caa caa agt gta ggt ggg ttc gat Ala Phe Ser Ala Val Ala Thr Cys Gln Gln Ser Val Gly Gly Phe Asp 680 685 690			2178
gat gct gac atg gat ggt gtt aac gtt ata agg ttt ggg aca aac cct Asp Ala Asp Met Asp Gly Val Asn Val Ile Arg Phe Gly Thr Asn Pro 695 700 705			2226
act ggt gac gtg tct ctc acg ctt ggt tta cgc cac gct gga aac atg Thr Gly Asp Val Ser Leu Thr Leu Gly Leu Arg His Ala Gly Asn Met 710 715 720			2274
cct gac aaa gac gct tct ttc tgc gtt aga gag ttt ggg ggt ttt tag Pro Asp Lys Asp Ala Ser Phe Cys Val Arg Glu Phe Gly Gly Phe 725 730 735			2322
tttgctttg tcactccatt taattaatta attatagttt tccattctta cttatTTAA ttgaaaatct attttgtct cttaaaagtc caaacaatac attagtctag ccctccctctg			2382
ctttttttt tctatctcgtaa gaagagaaga aaacgatacg taaatccctt cgaaaaactaa			2442
tgtacgttgc acgacttattt gtttcataa aaaaaaaaaaaa aaa			2502
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Phe His Asn Asn Ser Met Ser Gln Asp Tyr His His His His His His 20 25 30			
Asn Gln His Gln Gln Gly Ile Phe Asn Phe Ser Asn Gln Gly Phe Asp Arg 35 40 45			
Ser Asp Ser Pro Asn Leu Thr Thr Gln Gln Lys Gln Glu His Gln Arg			

MBI15 Sequence Listing .ST25

50                   55                   60

Val Glu Met Asp Glu Glu Ser Ser Val Ala Gly Gly Arg Ile Pro Val  
65                   70                   75                   80

Tyr Glu Ser Ala Gly Met Leu Ser Glu Met Phe Asn Phe Pro Gly Ser  
85                   90                   95

Ser Gly Gly Arg Asp Leu Asp Leu Gly Gln Ser Phe Arg Ser Asn  
100                  105                  110

Arg Gln Leu Leu Glu Glu Gln His Gln Asn Ile Pro Ala Met Asn Ala  
115                  120                  125

Thr Asp Ser Ala Thr Ala Ala Ala Met Gln Leu Phe Leu Met  
130                  135                  140

Asn Pro Pro Pro Pro Gln Gln Pro Pro Ser Pro Ser Ser Thr Thr Ser  
145                  150                  155                  160

Pro Arg Ser His His Asn Ser Ser Thr Leu His Met Leu Leu Pro Ser  
165                  170                  175

Pro Ser Thr Asn Thr His His Gln Asn Tyr Thr Asn His Met Ser  
180                  185                  190

Met His Gln Leu Pro His Gln His His Gln Gln Ile Ser Thr Trp Gln  
195                  200                  205

Ser Ser Pro Asp His His His His His Asn Ser Gln Thr Glu Ile  
210                  215                  220

Gly Thr Val His Val Glu Asn Ser Gly Gly His Gly Gln Gly Leu  
225                  230                  235                  240

Ser Leu Ser Leu Ser Ser Ser Leu Glu Ala Ala Ala Lys Ala Glu Glu  
245                  250                  255

Tyr Arg Asn Ile Tyr Tyr Gly Ala Asn Ser Ser Asn Ala Ser Pro His  
260                  265                  270

His Gln Tyr Asn Gln Phe Lys Thr Leu Leu Ala Asn Ser Ser Gln His  
275                  280                  285

His His Gln Val Leu Asn Gln Phe Arg Ser Ser Pro Ala Ala Ser Ser  
290                  295                  300

Ser Ser Met Ala Ala Val Asn Ile Leu Arg Asn Ser Arg Tyr Thr Thr  
305                  310                  315                  320

Ala Ala Gln Glu Leu Leu Glu Glu Phe Cys Ser Val Gly Arg Gly Phe  
325                  330                  335

Leu Lys Lys Asn Lys Leu Gly Asn Ser Ser Asn Pro Asn Thr Cys Gly  
340                  345                  350

MBI15 Sequence Listing.ST25

Gly Asp Gly Gly Ser Ser Pro Ser Ser Ala Gly Ala Asn Lys Glu  
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His Pro Pro Leu Ser Ala Ser Asp Arg Ile Glu His Gln Arg Arg Lys  
370 375 380

Val Lys Leu Leu Thr Met Leu Glu Glu Val Asp Arg Arg Tyr Asn His  
385 390 395 400

Tyr Cys Glu Gln Met Gln Met Val Val Asn Ser Phe Asp Ile Val Met  
405 410 415

Gly His Gly Ala Ala Leu Pro Tyr Thr Ala Leu Ala Gln Lys Ala Met  
420 425 430

Ser Arg His Phe Arg Cys Leu Lys Asp Ala Val Ala Ala Gln Leu Lys  
435 440 445

Gln Ser Cys Glu Leu Leu Gly Asp Lys Asp Ala Ala Gly Ile Ser Ser  
450 455 460

Ser Gly Leu Thr Lys Gly Glu Thr Pro Arg Leu Arg Leu Leu Glu Gln  
465 470 475 480

Ser Leu Arg Gln Gln Arg Ala Phe His Gln Met Gly Met Met Glu Gln  
485 490 495

Glu Ala Trp Arg Pro Gln Arg Gly Leu Pro Glu Arg Ser Val Asn Ile  
500 505 510

Leu Arg Ala Trp Leu Phe Glu His Phe Leu His Pro Tyr Pro Ser Asp  
515 520 525

Ala Asp Lys His Leu Leu Ala Arg Gln Thr Gly Leu Ser Arg Asn Gln  
530 535 540

Val Ser Asn Trp Phe Ile Asn Ala Arg Val Arg Leu Trp Lys Pro Met  
545 550 555 560

Val Glu Glu Met Tyr Gln Gln Glu Ser Lys Glu Arg Glu Arg Glu Glu  
565 570 575

Glu Leu Glu Glu Asn Glu Glu Asp Gln Glu Thr Lys Asn Ser Asn Asp  
580 585 590

Asp Lys Ser Thr Lys Ser Asn Asn Asn Glu Ser Asn Phe Thr Ala Val  
595 600 605

Arg Thr Thr Ser Gln Thr Pro Thr Thr Thr Ala Pro Asp Ala Ser Asp  
610 615 620

Ala Asp Ala Ala Val Ala Thr Gly His Arg Leu Arg Ser Asn Ile Asn  
625 630 635 640

Ala Tyr Glu Asn Asp Ala Ser Ser Leu Leu Leu Pro Ser Ser Tyr Ser  
645 650 655

## MBI15 Sequence Listing.ST25

Asn Ala Ala Ala Pro Ala Ala Val Ser Asp Asp Leu Asn Ser Arg Tyr  
 660 665 670

Gly Gly Ser Asp Ala Phe Ser Ala Val Ala Thr Cys Gln Gln Ser Val  
 675 680 685

Gly Gly Phe Asp Asp Ala Asp Met Asp Gly Val Asn Val Ile Arg Phe  
 690 695 700

Gly Thr Asn Pro Thr Gly Asp Val Ser Leu Thr Leu Gly Leu Arg His  
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Gly Gly Phe

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<223> G591

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1 5	
ctt tct gac caa act cct tct gat gat ttc ttc gag caa atc ctc ggc Leu Ser Asp Gln Thr Pro Ser Asp Asp Phe Glu Gln Ile Leu Gly	162
10 15 20 25	
ctt cct aac ttc tca gcc tct tct gcc gcc ggt tta tct gga gtt gac Leu Pro Asn Phe Ser Ala Ser Ser Ala Ala Gly Leu Ser Gly Val Asp	210
30 35 40	
gga gga tta ggt ggt gga gca ccg cct atg atg ctg cag ttg ggt tcc Gly Gly Leu Gly Gly Ala Pro Pro Met Met Leu Gln Leu Gly Ser	258
45 50 55	
gga gaa gaa gga agt cac atg ggt ggc tta gga gga agt gga cca act Gly Glu Gly Ser His Met Gly Leu Gly Gly Ser Gly Pro Thr	306
60 65 70	
ggg ttt cac aat cag atg ttt cct ttg ggg tta agt ctt gat caa ggg Gly Phe His Asn Gln Met Phe Pro Leu Gly Leu Ser Leu Asp Gln Gly	354
75 80 85	
aaa gga cct ggg ttt ctt aga cct gaa gga gga cat gga agt ggg aaa Lys Gly Pro Gly Phe Leu Arg Pro Glu Gly Gly His Gly Ser Gly Lys	402
90 95 100 105	
aga ttc tca gat gat gtt gat aat cga tgt tct tct atg aaa cct Arg Phe Ser Asp Asp Val Val Asp Asn Arg Cys Ser Ser Met Lys Pro	450
110 115 120	
gtt ttc cac ggg cag cct atg caa cag cca cct cca tcg gcc cca cat Val Phe His Gly Gln Pro Met Gln Gln Pro Pro Pro Ser Ala Pro His	498
125 130 135	

MBI15 Sequence Listing.ST25

cag cct act tca atc cgt ccc agg gtt cga gct agg cgt ggt cag gct Gln Pro Thr Ser Ile Arg Pro Arg Val Arg Ala Arg Arg Gly Gln Ala 140 145 150	546
act gat cca cat agc atc gct gag cgg cta cgt aga gaa aga ata gca Thr Asp Pro His Ser Ile Ala Glu Arg Leu Arg Arg Glu Arg Ile Ala 155 160 165	594
gaa cg <sup>g</sup> atc agg gcg ctg cag gaa ctt gta cct act gtg aac aag acc Glu Arg Ile Arg Ala Leu Gln Glu Leu Val Pro Thr Val Asn Lys Thr 170 175 180 185	642
gat aga gct gct atg atc gat gag att gtc gat tat gta aag ttt ctc Asp Arg Ala Ala Met Ile Asp Glu Ile Val Asp Tyr Val Lys Phe Leu 190 195 200	690
agg ctc caa gtc aag gtt ttg agc atg aac cga ctt ggt gga gcc ggt Arg Leu Gln Val Lys Val Leu Ser Met Asn Arg Leu Gly Gly Ala Gly 205 210 215	738
gcg gtt gct cca ctt gtt act gat atg cct ctt tca tca tca gtt gag Ala Val Ala Pro Leu Val Thr Asp Met Pro Leu Ser Ser Ser Val Glu 220 225 230	786
gat gaa acg ggt gag ggt gga agg act ccg caa cca gcg tgg gag aaa Asp Glu Thr Gly Glu Gly Arg Thr Pro Gln Pro Ala Trp Glu Lys 235 240 245	834
tgg tct aac gat ggg act gaa cgt caa gtg gct aaa ctg atg gaa gag Trp Ser Asn Asp Gly Thr Glu Arg Gln Val Ala Lys Leu Met Glu Glu 250 255 260 265	882
aac gtt gga gcc gcg atg cag ctt ctt caa tca aag gct ctt tgt atg Asn Val Gly Ala Ala Met Gln Leu Leu Gln Ser Lys Ala Leu Cys Met 270 275 280	930
atg cca atc tca ttg gca atg gca att tac cat tct caa cct ccg gat Met Pro Ile Ser Leu Ala Met Ala Ile Tyr His Ser Gln Pro Pro Asp 285 290 295	978
aca tct tca gtg gtc aag cct gag aac aat cct cca cag tag Thr Ser Ser Val Val Lys Pro Glu Asn Asn Pro Pro Gln 300 305 310	1020
gatttctgca ataaaagagtt tgtagacta atccaactgt ccaacatggg tttttttttt gctctaataatgc ctctggtttc ttctcttc tctcacccgac ttgaaaggtt aaaaagtgaa	1080 1140
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Asp Asp Phe Phe Glu Gln Ile Leu Gly Leu Pro Asn Phe Ser Ala Ser 20 25 30	
Ser Ala Ala Gly Leu Ser Gly Val Asp Gly Gly Leu Gly Gly Ala 35 40 45	
Pro Pro Met Met Leu Gln Leu Gly Ser Gly Glu Glu Gly Ser His Met 50 55 60	

## MBI15 Sequence Listing.ST25

Gly Gly Leu Gly Gly Ser Gly Pro Thr Gly Phe His Asn Gln Met Phe  
65 70 75 80

Pro Leu Gly Leu Ser Leu Asp Gln Gly Lys Gly Pro Gly Phe Leu Arg  
85 90 95

Pro Glu Gly Gly His Gly Ser Gly Lys Arg Phe Ser Asp Asp Val Val  
100 105 110

Asp Asn Arg Cys Ser Ser Met Lys Pro Val Phe His Gly Gln Pro Met  
115 120 125

Gln Gln Pro Pro Pro Ser Ala Pro His Gln Pro Thr Ser Ile Arg Pro  
130 135 140

Arg Val Arg Ala Arg Arg Gly Gln Ala Thr Asp Pro His Ser Ile Ala  
145 150 155 160

Glu Arg Leu Arg Arg Glu Arg Ile Ala Glu Arg Ile Arg Ala Leu Gln  
165 170 175

Glu Leu Val Pro Thr Val Asn Lys Thr Asp Arg Ala Ala Met Ile Asp  
180 185 190

Glu Ile Val Asp Tyr Val Lys Phe Leu Arg Leu Gln Val Lys Val Leu  
195 200 205

Ser Met Asn Arg Leu Gly Gly Ala Gly Ala Val Ala Pro Leu Val Thr  
210 215 220

Asp Met Pro Leu Ser Ser Val Glu Asp Glu Thr Gly Glu Gly Gly  
225 230 235 240

Arg Thr Pro Gln Pro Ala Trp Glu Lys Trp Ser Asn Asp Gly Thr Glu  
245 250 255

Arg Gln Val Ala Lys Leu Met Glu Glu Asn Val Gly Ala Ala Met Gln  
260 265 270

Leu Leu Gln Ser Lys Ala Leu Cys Met Met Pro Ile Ser Leu Ala Met  
275 280 285

Ala Ile Tyr His Ser Gln Pro Pro Asp Thr Ser Ser Val Val Lys Pro  
290 295 300

Glu Asn Asn Pro Pro Gln  
305 310

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<223> G525

## MBI15 Sequence Listing.ST25

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					Met Asp Tyr	
					1	
gag gca tca aga atc gtc gaa atg gta gaa gat gaa gaa cat ata gat						165
Glu Ala Ser Arg Ile Val Glu Met Val Glu Asp Glu Glu His Ile Asp						
5	10		15			
cta cca cca gga ttc aga ttt cac cct act gat gaa gaa ctc ata act						213
Leu Pro Pro Gly Phe Arg Phe His Pro Thr Asp Glu Glu Leu Ile Thr						
20	25		30		35	
cac tac ctc aaa cca aag gtt ttc aac act ttc ttc tct gct act gcc						261
His Tyr Leu Lys Pro Lys Val Phe Asn Thr Phe Phe Ser Ala Thr Ala						
40	45		50			
att ggt gaa gtt gat ctc aac aag att gag cct tgg gac tta cca tgg						309
Ile Gly Glu Val Asp Leu Asn Lys Ile Glu Pro Trp Asp Leu Pro Trp						
55	60		65			
aag gct aag atg gga gaa aaa gaa tgg tat ttc ttc tgt gtg aga gac						357
Lys Ala Lys Met Gly Glu Lys Glu Trp Tyr Phe Phe Cys Val Arg Asp						
70	75		80			
cgg aaa tac ccg acc ggt tta agg aca aac cgg gcg aca gaa gcc ggt						405
Arg Lys Tyr Pro Thr Gly Leu Arg Thr Asn Arg Ala Thr Glu Ala Gly						
85	90		95			
tat tgg aaa gcc aca gga aaa gac aaa gag ata ttc aag gga aaa tca						453
Tyr Trp Lys Ala Thr Gly Lys Asp Lys Glu Ile Phe Lys Gly Lys Ser						
100	105		110		115	
ctt gtg ggt atg aag aaa act ttg gtt ttc tat aaa gga aga gct cct						501
Leu Val Gly Met Lys Lys Thr Leu Val Phe Tyr Lys Gly Arg Ala Pro						
120	125		130			
aaa gga gtt aaa acc aat tgg gtt atg cat gaa tat cgt tta gaa ggc						549
Lys Gly Val Lys Thr Asn Trp Val Met His Glu Tyr Arg Leu Glu Gly						
135	140		145			
aaa tat tgt att gaa aat ctt ccc caa aca gct aag aac gaa tgg gtt						597
Lys Tyr Cys Ile Glu Asn Leu Pro Gln Thr Ala Lys Asn Glu Trp Val						
150	155		160			
ata tgt cgt gtt ttc caa aaa cgt gcc gat ggt aca aag gtt cca atg						645
Ile Cys Arg Val Phe Gln Lys Arg Ala Asp Gly Thr Lys Val Pro Met						
165	170		175			
tca atg ctt gat cca cac att aac cga atg gaa cca gcc ggt tta cct						693
Ser Met Leu Asp Pro His Ile Asn Arg Met Glu Pro Ala Gly Leu Pro						
180	185		190		195	
tcg tta atg gat tgt tct caa cga gac tcc ttc acc ggt tcg tcg tct						741
Ser Leu Met Asp Cys Ser Gln Arg Asp Ser Phe Thr Gly Ser Ser Ser						
200	205		210			
cac gtg acc tgc ttc tcc gac caa gaa acc gaa gac aaa aga ctt gtc						789
His Val Thr Cys Phe Ser Asp Gln Glu Thr Glu Asp Lys Arg Leu Val						
215	220		225			
cac gag tcc aaa gac ggt ttt ggt tct ctg ttt tac tcg gat cct ctg						837
His Glu Ser Lys Asp Gly Phe Gly Ser Leu Phe Tyr Ser Asp Pro Leu						
230	235		240			
ttt tta caa gac aat tat tcg cta atg aag ctg ttg ctt gac ggt caa						885
Phe Leu Gln Asp Asn Tyr Ser Leu Met Lys Leu Leu Asp Gly Gln						
245	250		255			
gaa act caa ttc tcc ggc aaa cct ttc gac ggt cgt gat tcg tcc ggt						933
Glu Thr Gln Phe Ser Gly Lys Pro Phe Asp Gly Arg Asp Ser Ser Gly						
260	265		270		275	

## MBI15 Sequence Listing .ST25

aca gaa gaa ttg gat tgc gtt tgg aat ttc tga gttgtataag ttatgttta	986
Thr Glu Glu Leu Asp Cys Val Trp Asn Phe	
280	285
gactttagt agtcatgtgt tcgtgtgtgt gaatgaatat tcttgtaa ttttttta	1046
aaaaaggaga aaaaaatatg ctagaaagtc aattgctttt gttatgttagc attagtgttt	1106
tttatgtact caatagactt cctaattaaa taaaaatctt aatttatttg caaaaaaaaa	1166
aaaaaaaaaa aaa	1179
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His Ile Asp Leu Pro Pro Gly Phe Arg Phe His Pro Thr Asp Glu Glu	
20	25
30	
Leu Ile Thr His Tyr Leu Lys Pro Lys Val Phe Asn Thr Phe Phe Ser	
35	40
45	
Ala Thr Ala Ile Gly Glu Val Asp Leu Asn Lys Ile Glu Pro Trp Asp	
50	55
60	
Leu Pro Trp Lys Ala Lys Met Gly Glu Lys Glu Trp Tyr Phe Phe Cys	
65	70
75	80
Val Arg Asp Arg Lys Tyr Pro Thr Gly Leu Arg Thr Asn Arg Ala Thr	
85	90
95	
Glu Ala Gly Tyr Trp Lys Ala Thr Gly Lys Asp Lys Glu Ile Phe Lys	
100	105
110	
Gly Lys Ser Leu Val Gly Met Lys Lys Thr Leu Val Phe Tyr Lys Gly	
115	120
125	
Arg Ala Pro Lys Gly Val Lys Thr Asn Trp Val Met His Glu Tyr Arg	
130	135
140	
Leu Glu Gly Lys Tyr Cys Ile Glu Asn Leu Pro Gln Thr Ala Lys Asn	
145	150
155	160
Glu Trp Val Ile Cys Arg Val Phe Gln Lys Arg Ala Asp Gly Thr Lys	
165	170
175	
Val Pro Met Ser Met Leu Asp Pro His Ile Asn Arg Met Glu Pro Ala	
180	185
190	
Gly Leu Pro Ser Leu Met Asp Cys Ser Gln Arg Asp Ser Phe Thr Gly	
195	200
205	
Ser Ser Ser His Val Thr Cys Phe Ser Asp Gln Glu Thr Glu Asp Lys	
210	215
220	

## MBI15 Sequence Listing.ST25

Arg Leu Val His Glu Ser Lys Asp Gly Phe Gly Ser Leu Phe Tyr Ser  
 225                   230                   235                   240

Asp Pro Leu Phe Leu Gln Asp Asn Tyr Ser Leu Met Lys Leu Leu Leu  
 245                   250                   255

Asp Gly Gln Glu Thr Gln Phe Ser Gly Lys Pro Phe Asp Gly Arg Asp  
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Ser Ser Gly Thr Glu Glu Leu Asp Cys Val Trp Asn Phe  
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 Met  
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gcg ctc gag gct ctt aca tca cca aga tta gct tct ccg att cct cct         105  
 Ala Leu Glu Ala Leu Thr Ser Pro Arg Leu Ala Ser Pro Ile Pro Pro  
 5                   10                   15

tgg ttc gaa gat tct tca gtc ttc cat gga gtc gag cac tgg aca aag         153  
 Leu Phe Glu Asp Ser Ser Val Phe His Gly Val Glu His Trp Thr Lys  
 20                   25                   30

ggg aag cga tct aag aga tca aga tcc gat ttc cac cac caa aac ctc         201  
 Gly Lys Arg Ser Lys Arg Ser Arg Ser Asp Phe His His Gln Asn Leu  
 35                   40                   45

act gag gaa gag tat cta gct ttt tgc ctc atg ctt ctc gct cgc gac         249  
 Thr Glu Glu Tyr Leu Ala Phe Cys Leu Met Leu Leu Ala Arg Asp  
 50                   55                   60                   65

aac cgt cag cct cct cct ccg gcg gtg gag aag ttg agc tac aag         297  
 Asn Arg Gln Pro Pro Pro Pro Ala Val Glu Lys Leu Ser Tyr Lys  
 70                   75                   80

tgt agc gtc tgc gac aag acg ttc tct tct tac caa gct ctc ggt ggt         345  
 Cys Ser Val Cys Asp Lys Thr Phe Ser Ser Tyr Gln Ala Leu Gly Gly  
 85                   90                   95

cac aag gca agc cac cgt aag aac tta tca cag act ctc tcc ggc gga         393  
 His Lys Ala Ser His Arg Lys Asn Leu Ser Gln Thr Leu Ser Gly Gly  
 100                   105                   110

gga gat gat cat tca acc tcg tcg gcg aca acc aca tcc gcc gtg act         441  
 Gly Asp Asp His Ser Thr Ser Ser Ala Thr Thr Thr Ser Ala Val Thr  
 115                   120                   125

act gga agt ggg aaa tca cac gtt tgc acc atc tgt aac aag tct ttt         489  
 Thr Gly Ser Gly Lys Ser His Val Cys Thr Ile Cys Asn Lys Ser Phe  
 130                   135                   140                   145

cct tcc ggt caa gct ctc ggc gga cac aag cgg tgc cac tac gaa gga         537  
 Pro Ser Gly Gln Ala Leu Gly Gly His Lys Arg Cys His Tyr Glu Gly  
 150                   155                   160

aac aac aac atc aac act agt agc gtg tcc aac tcc gaa ggt gcg ggg         585

MBI15 Sequence Listing ST25

Asn Asn Asn Ile Asn Thr Ser Ser Val Ser Asn Ser Glu Gly Ala Gly			
165	170	175	
tcc act agc cac gtt agc agt agc cac cgt ggg ttt gac ctc aac atc		633	
Ser Thr Ser His Val Ser Ser His Arg Gly Phe Asp Leu Asn Ile			
180	185	190	
cct ccg atc cct gaa ttc tcg atg gtc aac gga gac gac gaa gtc atg		681	
Pro Pro Ile Pro Glu Phe Ser Met Val Asn Gly Asp Asp Glu Val Met			
195	200	205	
agc cct atg ccg gcg aag aag cct cgg ttt gac ttt ccg gtc aaa ctt		729	
Ser Pro Met Pro Ala Lys Lys Pro Arg Phe Asp Phe Pro Val Lys Leu			
210	215	220	225
caa ctt taa ggaaatttac ttagacgata agatttcggt tgtatactgt		778	
Gln Leu			
tgagagttgt gtaggaattt gttgactgta cataccaaat tggactttga ctgattccaa		838	
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Pro Leu Phe Glu Asp Ser Ser Val Phe His Gly Val Glu His Trp Thr			
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Lys Gly Lys Arg Ser Lys Arg Ser Arg Ser Asp Phe His His Gln Asn			
35	40	45	
Leu Thr Glu Glu Glu Tyr Leu Ala Phe Cys Leu Met Leu Leu Ala Arg			
50	55	60	
Asp Asn Arg Gln Pro Pro Pro Pro Ala Val Glu Lys Leu Ser Tyr			
65	70	75	80
Lys Cys Ser Val Cys Asp Lys Thr Phe Ser Ser Tyr Gln Ala Leu Gly			
85	90	95	
Gly His Lys Ala Ser His Arg Lys Asn Leu Ser Gln Thr Leu Ser Gly			
100	105	110	
Gly Gly Asp Asp His Ser Thr Ser Ser Ala Thr Thr Thr Ser Ala Val			
115	120	125	
Thr Thr Gly Ser Gly Lys Ser His Val Cys Thr Ile Cys Asn Lys Ser			
130	135	140	
Phe Pro Ser Gly Gln Ala Leu Gly Gly His Lys Arg Cys His Tyr Glu			
145	150	155	160
Gly Asn Asn Asn Ile Asn Thr Ser Ser Val Ser Asn Ser Glu Gly Ala			
165	170	175	

MBI15 Sequence Listing .ST25

Gly	Ser	Thr	Ser	His	Val	Ser	Ser	Ser	His	Arg	Gly	Phe	Asp	Leu	Asn
180						185								190	

Ile	Pro	Pro	Ile	Pro	Glu	Phe	Ser	Met	Val	Asn	Gly	Asp	Asp	Glu	Val
195							200							205	

Met	Ser	Pro	Met	Pro	Ala	Lys	Lys	Pro	Arg	Phe	Asp	Phe	Pro	Val	Lys
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Leu Gln Leu  
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cacacctatt	attctcttgg	tgtgtttgtg	tgttacatat	acgtgtgagt	acataactttg										180	
ttgtaaaagt	ggatcgagg	tatggaaagg	gaccggttcc	accggaaaca	tcggcggcgg										240	
cggatgataa	ttcgtcttgg	aacgagactg	atgtcaccgc	c atg gtc tcc	gct ctc										296	
				Met	Val	Ser	Ala	Leu								
				1				5								
agc cgt gtc	ata gag aat	ccg aca gac	ccg ccg	gtc aaa	caa gag	ctt									344	
Ser Arg Val Ile	Glu Asn Pro Thr	Asp Pro Val Lys	Gln Glu Leu													
10	15	20														
gat aaa tcg	aat caa cat	caa cca gac	caa gat	caa cca	aga aga aga	aga									392	
Asp Lys Ser Asp	Gln His Gln Pro	Asp Gln Asp Gln	Asp Gln Pro Arg	Arg Arg Arg												
25	30	35														
cac tat aga	ggc gta agg	cag aga cca	tgg ggt	aaa tgg	gcg gca gaa										440	
His Tyr Arg Gly	Val Arg Gln Arg	Pro Trp Gly Lys	Trp Ala Ala Glu													
40	45	50														
atc cgc gat	cca aag aaa	gca gcc cgt	gtc tgg ctc	ggg act ttc	gag										488	
Ile Arg Asp Pro	Lys Ala Ala Arg	Val Trp Leu	Gly Thr Phe Glu													
55	60	65														
acg gca gag	gaa gct gct	tta gcc tat	gac cga	gct gcc	ctc aaa ttc										536	
Thr Ala Glu	Glu Ala Ala	Leu Ala Tyr	Asp Arg Ala	Ala Leu Lys	Phe											
70	75	80	85													
aaa ggc acc	aag gct aaa	ctg aac ttc	cct gaa	cg	gtc caa ggc cct										584	
Lys Gly Thr	Lys Ala Lys	Leu Asn Phe	Pro Glu Arg	Val Gln Gly	Pro											
90	95	100														
act acc acc	aca acc att	tct cat	gca cca	aga gga	gtt agt gaa tcc										632	
Thr Thr Thr	Thr Ile Ser His	Ala Pro Arg	Gly Val Ser	Glu Ser												
105	110	115														
atg aac tca	cct cct cga	cct ggt	cca cct tca	act act	act act										680	
Met Asn Ser	Pro Pro Arg	Pro Gly	Pro Pro Ser	Thr Thr	Thr Thr											
120	125	130														
tcg tgg cca	atg act tat	aac cag	gac ata	ctt caa	tac gct	cag ttg									728	
Ser Trp Pro	Met Thr Tyr	Asn Gln Asp	Ile Leu Gln	Tyr Ala Gln	Leu											
135	140	145														

## MBI15 Sequence Listing .ST25

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cag acg cag caa cag cag cta caa caa caa cag cag cgt gaa gaa Gln Thr Gln Gln Gln Leu Gln Gln Gln Gln Gln Arg Glu Glu 185 190 195	872
gaa gag aag aat tat ggt tac aat tat tat aac tac cca aga gaa taa Glu Glu Lys Asn Tyr Gly Tyr Asn Tyr Asn Tyr Pro Arg Glu 200 205 210	920
tctaattatt attgttggtc gaatcagttt tataaatagc tatcatagtt tcattttgg tttccgttaac ctttgttgcg tggaaaatat gaatgaacga gggacatgtg taacaatttg tttgtgttgcg taaaatgtta gttgtatttg gatttgctga agtttgattt tctgagcata aatcatttgcg cggtaaaaaa aaaaaaa	980 1040 1100 1126
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Gln Pro Arg Arg Arg His Tyr Arg Gly Val Arg Gln Arg Pro Trp Gly 35 40 45	
Lys Trp Ala Ala Glu Ile Arg Asp Pro Lys Lys Ala Ala Arg Val Trp 50 55 60	
Leu Gly Thr Phe Glu Thr Ala Glu Glu Ala Ala Leu Ala Tyr Asp Arg 65 70 75 80	
Ala Ala Leu Lys Phe Lys Gly Thr Lys Ala Lys Leu Asn Phe Pro Glu 85 90 95	
Arg Val Gln Gly Pro Thr Thr Thr Thr Ile Ser His Ala Pro Arg 100 105 110	
Gly Val Ser Glu Ser Met Asn Ser Pro Pro Pro Arg Pro Gly Pro Pro 115 120 125	
Ser Thr Thr Thr Ser Trp Pro Met Thr Tyr Asn Gln Asp Ile Leu 130 135 140	
Gln Tyr Ala Gln Leu Leu Thr Ser Asn Asn Glu Val Asp Leu Ser Tyr 145 150 155 160	
Tyr Thr Ser Thr Leu Phe Ser Gln Pro Phe Ser Thr Pro Ser Ser Ser 165 170 175	

## MBI15 Sequence Listing .ST25

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Gln Gln Arg Glu Glu Glu Lys Asn Tyr Gly Tyr Tyr Asn Tyr Tyr Asn  
 195 200 205

Tyr Pro Arg Glu  
 210

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Met Asp Gly Ser Ser Phe Leu Asp Ile Ser Leu Asp	
1 5 10	
ctc aac acc aat cct ttc tcc gca aaa ctt ccg aag aag gag gtc tca	159
Leu Asn Thr Asn Pro Phe Ser Ala Lys Leu Pro Lys Lys Glu Val Ser	
15 20 25	
gtt ttg gct tct act cac tta aag agg aaa tgg ttg gag caa gac gag	207
Val Leu Ala Ser Thr His Leu Lys Arg Lys Trp Leu Glu Gln Asp Glu	
30 35 40	
agc gca agt gag tta cga gag gag cta aac aga gtt aat tca gag aac	255
Ser Ala Ser Glu Leu Arg Glu Leu Asn Arg Val Asn Ser Glu Asn	
45 50 55 60	
aag aag cta aca gag atg tta gct aga gtc tgt gag agc tac aac gaa	303
Lys Lys Leu Thr Glu Met Leu Ala Arg Val Cys Glu Ser Tyr Asn Glu	
65 70 75	
cta cat aat cat ttg gag aag ctt cag agt cgc cag agc cct gaa atc	351
Leu His Asn His Leu Glu Lys Leu Gln Ser Arg Gln Ser Pro Glu Ile	
80 85 90	
gag cag acc gat ata ccg ata aag aaa aga aaa caa gac ccg gat gag	399
Glu Gln Thr Asp Ile Pro Ile Lys Lys Arg Lys Gln Asp Pro Asp Glu	
95 100 105	
ttc tta ggc ttt cct att gga ctc agt agt gga aaa act gag aac agc	447
Phe Leu Gly Phe Pro Ile Gly Leu Ser Ser Gly Lys Thr Glu Asn Ser	
110 115 120	
tcc agc aac gaa gat cat cat cat cat cag caa cat gag cag aaa	495
Ser Ser Asn Glu Asp His His His His Gln Gln His Glu Gln Lys	
125 130 135 140	
aat cag ctt ctt tca tgt aaa aga cca gtc act gat agc ttc aac aaa	543
Asn Gln Leu Leu Ser Cys Lys Arg Pro Val Thr Asp Ser Phe Asn Lys	
145 150 155	
gca aaa gtt tcg act gtc tac gtg cct act gaa aca tcg gac aca agc	591
Ala Lys Val Ser Thr Val Tyr Val Pro Thr Glu Thr Ser Asp Thr Ser	
160 165 170	
ttg aca gtt aaa gat gga ttt caa tgg agg aaa tac gga caa aag gtt	639
Leu Thr Val Lys Asp Gly Phe Gln Trp Arg Lys Tyr Gly Gln Lys Val	
175 180 185	

## MBI15 Sequence Listing.ST25

aca aga gac aac ccg tca cct aga gct tac ttt aga tgc tcg ttt gca Thr Arg Asp Asn Pro Ser Pro Arg Ala Tyr Phe Arg Cys Ser Phe Ala 190 195 200	687
ccg tct tgt cca gta aaa aag aag gta caa cgc agc gca gag gat cca Pro Ser Cys Pro Val Lys Lys Val Gln Arg Ser Ala Glu Asp Pro 205 210 215 220	735
tct tta ctt gta gcg aca tac gaa ggg acg cat aac cac ttg ggt cca Ser Leu Leu Val Ala Thr Tyr Glu Gly Thr His Asn His Leu Gly Pro 225 230 235	783
aat gct tct gaa ggg gat gct aca agc cag ggt ggg tca agc aca gtg Asn Ala Ser Glu Gly Asp Ala Thr Ser Gln Gly Ser Ser Thr Val 240 245 250	831
act ttg gat ctg gtt aat ggc tgt cat aga cta gcg ttg gag aaa aac Thr Leu Asp Leu Val Asn Gly Cys His Arg Leu Ala Leu Glu Lys Asn 255 260 265	879
gaa agg gat aat acg atg caa gag gtt ctg att caa caa atg gcg tca Glu Arg Asp Asn Thr Met Gln Glu Val Leu Ile Gln Gln Met Ala Ser 270 275 280	927
tcg tta aca aaa gat tcg aaa ttt aca gct gct ctt gct gct gct ata Ser Leu Thr Lys Asp Ser Lys Phe Thr Ala Ala Leu Ala Ala Ile 285 290 295 300	975
tct ggg agg tta atg gag caa tct aga aca tga acgttttag tgaatgtatt Ser Gly Arg Leu Met Glu Gln Ser Arg Thr 305 310	1028
gtttttgtt gtttagaatg atttttcgtt ttcaattgt gtctttcgat taggagataa aagatgtata taaaatattat aagtagatga agaaatcgta taagtaaaaa aaaaaaaaaa aaaaa	1088 1148 1152
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Thr His Leu Lys Arg Lys Trp Leu Glu Gln Asp Glu Ser Ala Ser Glu 35 40 45	
Leu Arg Glu Glu Leu Asn Arg Val Asn Ser Glu Asn Lys Lys Leu Thr 50 55 60	
Glu Met Leu Ala Arg Val Cys Glu Ser Tyr Asn Glu Leu His Asn His 65 70 75 80	
Leu Glu Lys Leu Gln Ser Arg Gln Ser Pro Glu Ile Glu Gln Thr Asp 85 90 95	
Ile Pro Ile Lys Lys Arg Lys Gln Asp Pro Asp Glu Phe Leu Gly Phe 100 105 110	

MBI15 Sequence Listing.ST25

Pro Ile Gly Leu Ser Ser Gly	115	Lys Thr Glu Asn Ser Ser Ser Asn Glu	120	125
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Asp His His His His Gln Gln His Glu Gln Lys Asn Gln Leu Leu	130	135	140
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Ser Cys Lys Arg Pro Val Thr Asp Ser Phe Asn Lys Ala Lys Val Ser	145	150	155	160
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Thr Val Tyr Val Pro Thr Glu Thr Ser Asp Thr Ser Leu Thr Val Lys	165	170	175
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Asp Gly Phe Gln Trp Arg Lys Tyr Gly Gln Lys Val Thr Arg Asp Asn	180	185	190
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Pro Ser Pro Arg Ala Tyr Phe Arg Cys Ser Phe Ala Pro Ser Cys Pro	195	200	205
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Val Lys Lys Lys Val Gln Arg Ser Ala Glu Asp Pro Ser Leu Leu Val	210	215	220
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Ala Thr Tyr Glu Gly Thr His Asn His Leu Gly Pro Asn Ala Ser Glu	225	230	235	240
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Gly Asp Ala Thr Ser Gln Gly Gly Ser Ser Thr Val Thr Leu Asp Leu	245	250	255
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Val Asn Gly Cys His Arg Leu Ala Leu Glu Lys Asn Glu Arg Asp Asn	260	265	270
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Thr Met Gln Glu Val Leu Ile Gln Gln Met Ala Ser Ser Leu Thr Lys	275	280	285
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Asp Ser Lys Phe Thr Ala Ala Leu Ala Ala Ile Ser Gly Arg Leu	290	295	300
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Met Glu Gln Ser Arg Thr	305	310
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Pro Pro Pro Ser Ser Ile Tyr Ala Pro Pro Met Leu Val Asn Cys Ser	
5 10 15	

gg tgc cgg acg cct ctc cag ctc cca tcc ggc gcc cga tct att cgc	151
Gly Cys Arg Thr Pro Leu Gln Leu Pro Ser Gly Ala Arg Ser Ile Arg	
20 25 30 35	

## MBI15 Sequence Listing.ST25

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cct cct ccg caa cct tcc tcc gcc cct tct ccg cct ccc caa atc cac Pro Pro Pro Gln Pro Ser Ser Ala Pro Ser Pro Pro Gln Ile His 55 60 65	247
gcg cct ccc ggt cag ctg cct cac ccc cat ggc agg aag agg gcc gtg Ala Pro Pro Gly Gln Leu Pro His Pro His Gly Arg Lys Arg Ala Val 70 75 80	295
atc tgt ggc atc tcg tat cgt ttc tct cgc cac gag ctc aaa ggc tgc Ile Cys Gly Ile Ser Tyr Arg Phe Ser Arg His Glu Leu Lys Gly Cys 85 90 95	343
atc aac gac gcc aag tgc atg cgt cac ctt ctc atc aac aaa ttc aaa Ile Asn Asp Ala Lys Cys Met Arg His Leu Leu Ile Asn Lys Phe Lys 100 105 110 115	391
ttc tcc cca gat tca att ctc atg ctt acc gag gaa act gat cca Phe Ser Pro Asp Ser Ile Leu Met Leu Thr Glu Glu Thr Asp Pro 120 125 130	439
tat cgt atc ccg acc aag caa aac atg agg atg gca ttg tat tgg ctc Tyr Arg Ile Pro Thr Lys Gln Asn Met Arg Met Ala Leu Tyr Trp Leu 135 140 145	487
gta cag gga tgc aca gca ggc gac tca ctt gtc ttc cac tac tct ggt Val Gln Gly Cys Thr Ala Gly Asp Ser Leu Val Phe His Tyr Ser Gly 150 155 160	535
cat ggt tcg cgt caa aga aac tac aac ggt gat gaa gtt gat ggc tat His Gly Ser Arg Gln Arg Asn Tyr Asn Gly Asp Glu Val Asp Gly Tyr 165 170 175	583
gat gaa aca ctc tgt cct ctg gat ttt gaa act cag ggg atg att gta Asp Glu Thr Leu Cys Pro Leu Asp Phe Glu Thr Gln Gly Met Ile Val 180 185 190 195	631
gac gat gag atc aac gca acc att gta cgc cct ctt cca cat ggt gtc Asp Asp Glu Ile Asn Ala Thr Ile Val Arg Pro Leu Pro His Gly Val 200 205 210	679
aag ctc cat tca att atc gat gct tgc cat agt ggt acc gtt ctg gat Lys Leu His Ser Ile Ile Asp Ala Cys His Ser Gly Thr Val Leu Asp 215 220 225	727
tta ccc ttc cta tgc aga atg aac aga gct ggg cag tat gtg tgg gag Leu Pro Phe Leu Cys Arg Met Asn Arg Ala Gly Gln Tyr Val Trp Glu 230 235 240	775
gat cat cgg cct agg tca ggt ttg tgg aaa gga act gct ggt gga gaa Asp His Arg Pro Arg Ser Gly Leu Trp Lys Gly Thr Ala Gly Glu 245 250 255	823
gcc att tca att agt gga tgt gat gat gat cag act tcg gcc gac aca Ala Ile Ser Ile Ser Gly Cys Asp Asp Gln Thr Ser Ala Asp Thr 260 265 270 275	871
tca gcg ctg tcg aag atc acg tct acg ggt gct atg act ttc tgt ttt Ser Ala Leu Ser Lys Ile Thr Ser Thr Gly Ala Met Thr Phe Cys Phe 280 285 290	919
att caa gca att gaa cgc agc gca caa ggc aca acc tat gga agc ctt Ile Gln Ala Ile Glu Arg Ser Ala Gln Gly Thr Thr Tyr Gly Ser Leu 295 300 305	967
ctg aat tct atg cgc acc aca ata agg aat aca ggg aat gat ggt ggt Leu Asn Ser Met Arg Thr Thr Ile Arg Asn Thr Gly Asn Asp Gly Gly 310 315 320	1015
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MBI15 Sequence Listing .ST25

325	330	335	
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caa aca ttc gat gtc tat gca aag cct ttc act ctc tag taaaggacaa Gln Thr Phe Asp Val Tyr Ala Lys Pro Phe Thr Leu 360               365			
gtcacttttt atgtatagcg agtgtgattt gagaatccgt ccatataacc accttttgtt      1220			
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Ser Ile Arg Cys Ala Leu Cys Gln Ala Val Thr His Ile Ala Asp Pro 35               40               45			
Arg Thr Ala Pro Pro Pro Gln Pro Ser Ser Ala Pro Ser Pro Pro Pro 50               55               60			
Gln Ile His Ala Pro Pro Gly Gln Leu Pro His Pro His Gly Arg Lys 65               70               75               80			
Arg Ala Val Ile Cys Gly Ile Ser Tyr Arg Phe Ser Arg His Glu Leu 85               90               95			
Lys Gly Cys Ile Asn Asp Ala Lys Cys Met Arg His Leu Leu Ile Asn 100               105               110			
Lys Phe Lys Phe Ser Pro Asp Ser Ile Leu Met Leu Thr Glu Glu Glu 115               120               125			
Thr Asp Pro Tyr Arg Ile Pro Thr Lys Gln Asn Met Arg Met Ala Leu 130               135               140			
Tyr Trp Leu Val Gln Gly Cys Thr Ala Gly Asp Ser Leu Val Phe His 145               150               155               160			
Tyr Ser Gly His Gly Ser Arg Gln Arg Asn Tyr Asn Gly Asp Glu Val 165               170               175			
Asp Gly Tyr Asp Glu Thr Leu Cys Pro Leu Asp Phe Glu Thr Gln Gly 180               185               190			
Met Ile Val Asp Asp Glu Ile Asn Ala Thr Ile Val Arg Pro Leu Pro 195               200               205			
His Gly Val Lys Leu His Ser Ile Ile Asp Ala Cys His Ser Gly Thr			

MBI15 Sequence Listing .ST25

210	215	220
Val Leu Asp Leu Pro Phe Leu Cys Arg Met Asn Arg Ala Gly Gln Tyr		
225	230	235 240
Val Trp Glu Asp His Arg Pro Arg Ser Gly Leu Trp Lys Gly Thr Ala		
245	250	255
Gly Gly Glu Ala Ile Ser Ile Ser Gly Cys Asp Asp Asp Gln Thr Ser		
260	265	270
Ala Asp Thr Ser Ala Leu Ser Lys Ile Thr Ser Thr Gly Ala Met Thr		
275	280	285
Phe Cys Phe Ile Gln Ala Ile Glu Arg Ser Ala Gln Gly Thr Thr Tyr		
290	295	300
Gly Ser Leu Leu Asn Ser Met Arg Thr Thr Ile Arg Asn Thr Gly Asn		
305	310	315 320
Asp Gly Gly Ser Gly Gly Val Val Thr Thr Val Leu Ser Met Leu		
325	330	335
Leu Thr Gly Gly Ser Ala Ile Gly Gly Leu Arg Gln Glu Pro Gln Leu		
340	345	350
Thr Ala Cys Gln Thr Phe Asp Val Tyr Ala Lys Pro Phe Thr Leu		
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Met Ala Ser Ser Ser Ser Ser Tyr Arg Phe Gln Ser Gly Ser Tyr		
1 5 10 15		
cct ctt tcg tca agt cct tct ggg aat ttc gtc gaa cgc att aaa	96	
Pro Leu Ser Ser Pro Ser Leu Gly Asn Phe Val Glu Arg Ile Lys		
20 25 30		
gac gct tgt cat ttc ctt gtc tct gct gtt ttg ggt acc att atc tcc	144	
Asp Ala Cys His Phe Leu Val Ser Ala Val Leu Gly Thr Ile Ile Ser		
35 40 45		
gcg atc ttg acc ttc ttc gca cta gtg ggc aca ttg cta ggg gca	192	
Ala Ile Leu Thr Phe Phe Ala Leu Val Gly Thr Leu Leu Gly Ala		
50 55 60		
ctt aca gga gct ttg ata ggt caa gaa act gag agt ggt ttc att aga	240	
Leu Thr Gly Ala Leu Ile Gly Gln Glu Thr Glu Ser Gly Phe Ile Arg		
65 70 75 80		
gga gca gca att gga gcc att tcg gga gct gtt ttc tct atc gag gtc	288	
Gly Ala Ala Ile Gly Ala Ile Ser Gly Ala Val Phe Ser Ile Glu Val		
85 90 95		

MBI15 Sequence Listing .ST25

ttt gaa tca tct ctg gat ctc tgg aaa tcc gat gag tcg ggt ttc gga Phe Glu Ser Ser Leu Asp Leu Trp Lys Ser Asp Glu Ser Gly Phe Gly 100 105 110	336
tgt ttt ctc tac ttg att gat gtc att gtt agt ctt cta agc ggg aga Cys Phe Leu Tyr Leu Ile Asp Val Ile Val Ser Leu Leu Ser Gly Arg 115 120 125	384
ctt gta cga gag cgc att ggt cct gca atg cta agt gca gtg caa agt Leu Val Arg Glu Arg Ile Gly Pro Ala Met Leu Ser Ala Val Gln Ser 130 135 140	432
caa atg gga gct gtg gat aca gct ttt gat gat cac aca agc ctt ttt Gln Met Gly Ala Val Asp Thr Ala Phe Asp Asp His Thr Ser Leu Phe 145 150 155 160	480
gat aca gga ggc tca aaa gga ttg aca gga gac ctt gtt gag aaa atc Asp Thr Gly Gly Ser Lys Gly Leu Thr Gly Asp Leu Val Glu Lys Ile 165 170 175	528
cca aag atg aca atc act ggc aac aat aac act gat gct tct gag aac Pro Lys Met Thr Ile Thr Gly Asn Asn Asn Thr Asp Ala Ser Glu Asn 180 185 190	576
aca gac tca tgt tct gtt tgt cat cac atg ttt cac tta cct tgc ata Thr Asp Ser Cys Ser Val Cys Leu Gln Asp Phe Gln Leu Gly Glu Thr 195 200 205	624
gtt aga agc ttg cct cat tgt cat cac atg ttt cac tta cct tgc ata Val Arg Ser Leu Pro His Cys His Met Phe His Leu Pro Cys Ile 210 215 220	672
gac aat tgg ctc ctt aga cac ggt tct tgc ccg atg tgt aga cgt gat Asp Asn Trp Leu Leu Arg His Gly Ser Cys Pro Met Cys Arg Arg Asp 225 230 235 240	720
att taa Ile	726
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Pro Leu Ser Ser Ser Pro Ser Leu Gly Asn Phe Val Glu Arg Ile Lys 20 25 30	
Asp Ala Cys His Phe Leu Val Ser Ala Val Leu Gly Thr Ile Ile Ser 35 40 45	
Ala Ile Leu Thr Phe Phe Ala Leu Val Gly Thr Leu Leu Gly Ala 50 55 60	
Leu Thr Gly Ala Leu Ile Gly Gln Glu Thr Glu Ser Gly Phe Ile Arg 65 70 75 80	
Gly Ala Ala Ile Gly Ala Ile Ser Gly Ala Val Phe Ser Ile Glu Val 85 90 95	
Phe Glu Ser Ser Leu Asp Leu Trp Lys Ser Asp Glu Ser Gly Phe Gly 100 105 110	

## MBI15 Sequence Listing .ST25

Cys Phe Leu Tyr Leu Ile Asp Val Ile Val Ser Leu Leu Ser Gly Arg  
 115                   120                   125

Leu Val Arg Glu Arg Ile Gly Pro Ala Met Leu Ser Ala Val Gln Ser  
 130                   135                   140

Gln Met Gly Ala Val Asp Thr Ala Phe Asp Asp His Thr Ser Leu Phe  
 145                   150                   155                   160

Asp Thr Gly Gly Ser Lys Gly Leu Thr Gly Asp Leu Val Glu Lys Ile  
 165                   170                   175

Pro Lys Met Thr Ile Thr Gly Asn Asn Asn Thr Asp Ala Ser Glu Asn  
 180                   185                   190

Thr Asp Ser Cys Ser Val Cys Leu Gln Asp Phe Gln Leu Gly Glu Thr  
 195                   200                   205

Val Arg Ser Leu Pro His Cys His His Met Phe His Leu Pro Cys Ile  
 210                   215                   220

Asp Asn Trp Leu Leu Arg His Gly Ser Cys Pro Met Cys Arg Arg Asp  
 225                   230                   235                   240

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cggcaagttt	gtatctagaa	aggatcgatt	ggtgaggatca	atagtggttg	gtgggtttta	180
gta atg gaa gac	ggt gag ctt	aat tcc aat cag	gaa gtg ttt tcg			228
Met Glu Asp Gly	Glu Leu Asp Phe	Ser Asn Gln Glu Val	Phe Ser			
1	5	10	15			
agt tcg gag atg	ggt gaa tta cca	cct agc aat tgt	tcg atg gat	agt		276
Ser Ser Glu Met	Gly Glu Leu Pro	Pro Ser Asn Cys	Ser Met Asp	Ser		
20	25		30			
ttc ttt gat	ggg ctt tta	atg gat act	aat gct gct	tgt acc cac act		324
Phe Phe Asp	Gly Leu Leu	Met Asp Thr	Asn Ala Ala	Cys Thr His		
35	40	45				
cac acc tgt aac	ccc act gga	cca gag aac	act cat	act cac acg	tgc	372
His Thr Cys Asn	Pro Thr Gly	Pro Glu Asn	Thr His	Thr His	Thr Cys	
50	55	60				
ttc cat gtc	cac acc aag	att ctc ccg	gat gag	agc gat	gaa aaa gtt	420
Phe His Val	His Thr Lys	Ile Leu Pro	Asp Glu Ser	Asp Glu	Lys Val	
65	70	75				

MBI15 Sequence Listing .ST25

tct act gat gat aca gct gag tct tgt ggg aag aag ggt gaa aag aga Ser Thr Asp Asp Thr Ala Glu Ser Cys Gly Lys Lys Gly Glu Lys Arg 80 85 90 95	468
cct ttg gga aac cgg gaa gcg gtt aga aag tat aga gag aag aag aag Pro Leu Gly Asn Arg Glu Ala Val Arg Lys Tyr Arg Glu Lys Lys Lys 100 105 110	516
gct aaa gct gct tct ttg gag gat gag gtt gca agg ctt agg gcg gtg Ala Lys Ala Ala Ser Leu Glu Asp Glu Val Ala Arg Leu Arg Ala Val 115 120 125	564
aat cag cag ctg gtg aag agg ttg caa aat cag gct acc ttg gaa gct Asn Gln Gln Leu Val Lys Arg Leu Gln Asn Gln Ala Thr Leu Glu Ala 130 135 140	612
gag gtt tcg agg ctt aag tgt ttg ctt gtg gat ttg aga gga aga ata Glu Val Ser Arg Leu Lys Cys Leu Leu Val Asp Leu Arg Gly Arg Ile 145 150 155	660
gat gga gag att gga tct ttg cct tat cag aaa cct atg gct gca aat Asp Gly Glu Ile Gly Ser Phe Pro Tyr Gln Lys Pro Met Ala Ala Asn 160 165 170 175	708
att cct tct ttc tcg cac atg atg aat cct tgt aat gta caa tgt gat Ile Pro Ser Phe Ser His Met Met Asn Pro Cys Asn Val Gln Cys Asp 180 185 190	756
gat gaa gtt tat tgc cct cag aat gtg ttt gga gtg aat agc caa gaa Asp Glu Val Tyr Cys Pro Gln Asn Val Phe Gly Val Asn Ser Gln Glu 195 200 205	804
ggg gcc tcg atc aat gac caa ggg tta agt ggt tgt gat ttt gat cag Gly Ala Ser Ile Asn Asp Gln Gly Leu Ser Gly Cys Asp Phe Asp Gln 210 215 220	852
cta caa tgc atg gct aat cag aac tta aat gga aat gga aac gga tca Leu Gln Cys Met Ala Asn Gln Asn Leu Asn Gly Asn Gly Asn Gly Ser 225 230 235	900
tcc agc aac gtc aataca tct gtc tcg aat aag aga aag ggt ggg cat Phe Ser Asn Val Asn Thr Ser Val Ser Asn Lys Arg Lys Gly Gly His 240 245 250 255	948
cgt gca tca aga gca gtt tga agcatcatca agcttgtact atctatttcc Arg Ala Ser Arg Ala Val 260	999
accaggatag atattgtatt ccaaataagt tgtagagttc agctgcaggag tcagcttcgc tcagcttga ggggttggtg gtgtggctt tctttgtggc acgagtgaga tctatggaca gaacccagat ttagtagtag tagaggcagg atttcgactt ccactaacca tcatgttgct tggtaagaa caaggtatgc ccatgaagca cactgtttt tacattgagc ttgaggggct gtctctgatc tagccttact gtaacattgc aacgttctca caattgtatcccagttgc tttgttact taaatgtat aatatagctt aacttttact tgaaaaaaaaaaaaaaaaaaaa aaaaaaaaaaa a	1059 1119 1179 1239 1299 1359 1370
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Ser Glu Met Gly Glu Leu Pro Pro Ser Asn Cys Ser Met Asp Ser Phe

MBI15 Sequence Listing.ST25  
20 25 30

Phe Asp Gly Leu Leu Met Asp Thr Asn Ala Ala Cys Thr His Thr His  
35 40 45

Thr Cys Asn Pro Thr Gly Pro Glu Asn Thr His Thr His Thr Cys Phe  
50 55 60

His Val His Thr Lys Ile Leu Pro Asp Glu Ser Asp Glu Lys Val Ser  
65 70 75 80

Thr Asp Asp Thr Ala Glu Ser Cys Gly Lys Lys Gly Glu Lys Arg Pro  
85 90 95

Leu Gly Asn Arg Glu Ala Val Arg Lys Tyr Arg Glu Lys Lys Ala  
100 105 110

Lys Ala Ala Ser Leu Glu Asp Glu Val Ala Arg Leu Arg Ala Val Asn  
115 120 125

Gln Gln Leu Val Lys Arg Leu Gln Asn Gln Ala Thr Leu Glu Ala Glu  
130 135 140

Val Ser Arg Leu Lys Cys Leu Leu Val Asp Leu Arg Gly Arg Ile Asp  
145 150 155 160

Gly Glu Ile Gly Ser Phe Pro Tyr Gln Lys Pro Met Ala Ala Asn Ile  
165 170 175

Pro Ser Phe Ser His Met Met Asn Pro Cys Asn Val Gln Cys Asp Asp  
180 185 190

Glu Val Tyr Cys Pro Gln Asn Val Phe Gly Val Asn Ser Gln Glu Gly  
195 200 205

Ala Ser Ile Asn Asp Gln Gly Leu Ser Gly Cys Asp Phe Asp Gln Leu  
210 215 220

Gln Cys Met Ala Asn Gln Asn Leu Asn Gly Asn Gly Asn Gly Ser Phe  
225 230 235 240

Ser Asn Val Asn Thr Ser Val Ser Asn Lys Arg Lys Gly Gly His Arg  
245 250 255

Ala Ser Arg Ala Val  
260

<210> 35  
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MBI15 Sequence Listing.ST25

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tttacttgtg caccttcaag atttcgtttt ttccagcgcc cagaatgctc cgggtgacca	120
acatttgttc ctgattcatt tcctattggc tcgtattgtc tgtcacaca agagaaattt	180
caagaagttg ttactaaaag agaggccaca agtggatatt gtctttgtta tcaagtgtta	240
gtacagaaaa gtggtgagaa agtaat atg gct gat acc agt ccg aga act gat Met Ala Asp Thr Ser Pro Arg Thr Asp	293
1 5	
gtc tca aca gat gac gac aca gat cat cct gat ctt ggg tcg gag gga Val Ser Thr Asp Asp Asp Thr Asp His Pro Asp Leu Gly Ser Glu Gly	341
10 15 20 25	
gca cta gtg aat act gct gct tct gat tcg agt gac cga tcg aag gga Ala Leu Val Asn Thr Ala Ala Ser Asp Ser Ser Asp Arg Ser Lys Gly	389
30 35 40	
aag atg gat caa aag act ctt cgt agg ctt gct caa aac cgt gag gca Lys Met Asp Gln Lys Thr Leu Arg Arg Leu Ala Gln Asn Arg Glu Ala	437
45 50 55	
gca agg aaa agc aga ttg agg aag aag gct tat gtt cag cag cta gag Ala Arg Lys Ser Arg Leu Arg Lys Lys Ala Tyr Val Gln Gln Leu Glu	485
60 65 70	
aac agc cgc ttg aaa cta acc cag ctt gag cag gag ctg caa aga gca Asn Ser Arg Leu Lys Leu Thr Gln Leu Glu Gln Glu Leu Gln Arg Ala	533
75 80 85	
aga cag cag ggc gtc ttc att tca ggc aca gga gac cag gcc cat tct Arg Gln Gln Gly Val Phe Ile Ser Gly Thr Gly Asp Gln Ala His Ser	581
90 95 100 105	
act ggt gga aat ggt gct ttg gcg ttt gat gct gaa cat tca cgg tgg Thr Gly Asn Gly Ala Leu Ala Phe Asp Ala Glu His Ser Arg Trp	629
110 115 120	
ttg gaa gaa aag aac aag caa atg aac gag ctg agg tct gct ctg aat Leu Glu Glu Lys Asn Lys Gln Met Asn Glu Leu Arg Ser Ala Leu Asn	677
125 130 135	
gcg cat gca ggt gat tct gag ctt cga ata ata gtc gat ggt gtg atg Ala His Ala Gly Asp Ser Glu Leu Arg Ile Ile Val Asp Gly Val Met	725
140 145 150	
gct cac tat gag gag ctt ttc agg ata aag agc aat gca gct aag aat Ala His Tyr Glu Glu Leu Phe Arg Ile Lys Ser Asn Ala Ala Lys Asn	773
155 160 165	
gat gtc ttt cac ttg cta tct ggc atg tgg aaa aca cca gct gag aga Asp Val Phe His Leu Leu Ser Gly Met Trp Lys Thr Pro Ala Glu Arg	821
170 175 180 185	
tgt ttc ttg tgg ctc ggt gga ttt cgt tca tcc gaa ctt cta aag ctt Cys Phe Leu Trp Leu Gly Gly Phe Arg Ser Ser Glu Leu Leu Lys Leu	869
190 195 200	
ctg gcg aat cag ttg gag cca atg aca gag aga cag ttg atg ggc ata Leu Ala Asn Gln Leu Glu Pro Met Thr Glu Arg Gln Leu Met Gly Ile	917
205 210 215	
aat aac ctg caa cag aca tcg cag cag gct gaa gat gct ttg tct caa Asn Asn Leu Gln Gln Thr Ser Gln Gln Ala Glu Asp Ala Leu Ser Gln	965
220 225 230	
ggg atg gag agc tta caa cag tca cta gct gat act tta tcg agc ggg Gly Met Glu Ser Leu Gln Gln Ser Leu Ala Asp Thr Leu Ser Ser Gly	1013
235 240 245	
act ctt ggt tca agt tca tca ggg aat gtc gca agc tac atg ggt cag Thr Leu Gly Ser Ser Ser Gly Asn Val Ala Ser Tyr Met Gly Gln	1061
250 255 260 265	

## MBI15 Sequence Listing.ST25

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cag gct gat aat ttg aga cta caa aca ttg caa cag atg ata aga gta Gln Ala Asp Asn Leu Arg Leu Gln Thr Leu Gln Gln Met Ile Arg Val 285 290 295	1157
tta aca acg aga cag tca gca cgt gct cta ctt gca ata cac gat tac Leu Thr Thr Arg Gln Ser Ala Arg Ala Leu Leu Ala Ile His Asp Tyr 300 305 310	1205
ttc tca cgg cta cga gct cta agc tcc tta tgg ctt gct cga ccc aga Phe Ser Arg Leu Arg Ala Leu Ser Ser Leu Trp Leu Ala Arg Pro Arg 315 320 325	1253
gag tga aactgtattt tggcacatg tcagctgtac aaaatccata tggcacacaaa Glu 330	1309
accaggagag actattaatc aacacttgtc agattcttct taccaaatcc atcaacaaat aagcaaattt ctggaaaca aaagactctt tgtatgttagg tttcttctac atggttgtgg taattcatgt tgtttagtt gtgtcatca gtttttaatt tagcatttga aaagttcaat gttggttata tagcatcttc gattatctta gaaaggttat tgaattttgt tttttttgt tacttttgtg tgtggtaaag gtgttttaac cttgcaactt ctgtactgtt atcatttaac aatattaaga tgttctattt gagttttgt	1369 1429 1489 1549 1609 1638
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Asp His Pro Asp Leu Gly Ser Glu Gly Ala Leu Val Asn Thr Ala Ala 20 25 30	
Ser Asp Ser Ser Asp Arg Ser Lys Gly Lys Met Asp Gln Lys Thr Leu 35 40 45	
Arg Arg Leu Ala Gln Asn Arg Glu Ala Ala Arg Lys Ser Arg Leu Arg 50 55 60	
Lys Lys Ala Tyr Val Gln Gln Leu Glu Asn Ser Arg Leu Lys Leu Thr 65 70 75 80	
Gln Leu Glu Gln Glu Leu Gln Arg Ala Arg Gln Gln Gly Val Phe Ile 85 90 95	
Ser Gly Thr Gly Asp Gln Ala His Ser Thr Gly Gly Asn Gly Ala Leu 100 105 110	
Ala Phe Asp Ala Glu His Ser Arg Trp Leu Glu Glu Lys Asn Lys Gln 115 120 125	
Met Asn Glu Leu Arg Ser Ala Leu Asn Ala His Ala Gly Asp Ser Glu 130 135 140	

## MBI15 Sequence Listing .ST25

Leu Arg Ile Ile Val Asp Gly Val Met Ala His Tyr Glu Glu Leu Phe  
 145                150                155                160

Arg Ile Lys Ser Asn Ala Ala Lys Asn Asp Val Phe His Leu Leu Ser  
 165                170                175

Gly Met Trp Lys Thr Pro Ala Glu Arg Cys Phe Leu Trp Leu Gly Gly  
 180                185                190

Phe Arg Ser Ser Glu Leu Leu Lys Leu Leu Ala Asn Gln Leu Glu Pro  
 195                200                205

Met Thr Glu Arg Gln Leu Met Gly Ile Asn Asn Leu Gln Gln Thr Ser  
 210                215                220

Gln Gln Ala Glu Asp Ala Leu Ser Gln Gly Met Glu Ser Leu Gln Gln  
 225                230                235                240

Ser Leu Ala Asp Thr Leu Ser Ser Gly Thr Leu Gly Ser Ser Ser Ser  
 245                250                255

Gly Asn Val Ala Ser Tyr Met Gly Gln Met Ala Met Ala Met Gly Lys  
 260                265                270

Leu Gly Thr Leu Glu Gly Phe Ile Arg Gln Ala Asp Asn Leu Arg Leu  
 275                280                285

Gln Thr Leu Gln Gln Met Ile Arg Val Leu Thr Thr Arg Gln Ser Ala  
 290                295                300

Arg Ala Leu Leu Ala Ile His Asp Tyr Phe Ser Arg Leu Arg Ala Leu  
 305                310                315                320

Ser Ser Leu Trp Leu Ala Arg Pro Arg Glu  
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 Met Asp Gly Glu Asp Phe Ala Gly Lys Ala  
 1                5                10

gct gct gaa gcc aag gga ttg aac ccg gga tta atc gtg ctg ctt gtt      160  
 Ala Ala Glu Ala Lys Gly Leu Asn Pro Gly Leu Ile Val Leu Val  
 15                20                25

gtt gga ggt ccg ctt ctt gtg ttc cta atc gcc aac tac gtg ctt tac      208  
 Val Gly Gly Pro Leu Leu Val Phe Leu Ile Ala Asn Tyr Val Leu Tyr  
 30                35                40

## MBI15 Sequence Listing .ST25

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aaa aag aag ctc aag cgg gag aag cta aag caa gga gtc cct gtc cct Lys Lys Lys Leu Lys Arg Glu Lys Leu Lys Gln Gly Val Pro Val Pro 60 65 70	304
gga gaa taa aaggcagctt aagtttcctt cacttgcgc tccttcaaag Gly Glu 75	353
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Val Phe Leu Ile Ala Asn Tyr Val Leu Tyr Val Tyr Ala Gln Lys Asn 35 40 45	
Leu Pro Pro Arg Lys Lys Pro Val Ser Lys Lys Lys Leu Lys Arg 50 55 60	
Glu Lys Leu Lys Gln Gly Val Pro Val Pro Gly Glu 65 70 75	
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aaatcaaaga gactttgaa gattgttcc caatttgcgt caatcgggat cgagtcaaat 180	
ctgaaatctt ctccactcat catctgacta taagacttaa tcaagggact ttttggcg 240	
gtttggtttt aaacgtcttg gattcgaagt ggtaaggt atg gat gaa aat aat Met Asp Glu Asn Asn 1 5	294
gga ggt tca agc tca ctt cca cct ttc ctt act aaa aca tat gaa atg Gly Gly Ser Ser Ser Leu Pro Pro Phe Leu Thr Lys Thr Tyr Glu Met 10 15 20	342
gtt gat gat tct tct gac tcg gtc gtt gct tgg agc gaa aac aac Val Asp Asp Ser Ser Asp Ser Val Val Ala Trp Ser Glu Asn Asn	390

MBI15 Sequence Listing.ST25

25	30	35	
aaa agc ttc atc gtc aag aat cca gca gag ttt tca aga gac ctt ctt			438
Lys Ser Phe Ile Val Lys Asn Pro Ala Glu Phe Ser Arg Asp Leu Leu			
40	45	50	
ccg aga ttc ttc aag cat aag aat ttc tca agt ttc atc cgt cag ctt			486
Pro Arg Phe Phe Lys His Lys Asn Phe Ser Ser Phe Ile Arg Gln Leu			
55	60	65	
aat aca tat ggt ttt cga aaa gta gat cct gag aaa tgg gaa ttc ttg			534
Asn Thr Tyr Gly Phe Arg Lys Val Asp Pro Glu Lys Trp Glu Phe Leu			
70	75	80	85
aat gat gat ttt gtt aga ggt cga cct tac ctt atg aag aac att cat			582
Asn Asp Asp Phe Val Arg Gly Arg Pro Tyr Leu Met Lys Asn Ile His			
90	95	100	
aga cga aaa ccg gtt cat agc cac tcg tta gtg aat cta caa gcg caa			630
Arg Arg Lys Pro Val His Ser His Ser Leu Val Asn Leu Gln Ala Gln			
105	110	115	
aat cct ttg acg gaa tca gaa aga cgg agc atg gag gat cag ata gaa			678
Asn Pro Leu Thr Glu Ser Glu Arg Arg Ser Met Glu Asp Gln Ile Glu			
120	125	130	
aga ctg aaa aat gag aaa gaa ggc ctt ctt gcg gag tta cag aac caa			726
Arg Leu Lys Asn Glu Lys Glu Gly Leu Leu Ala Glu Leu Gln Asn Gln			
135	140	145	
gag caa gaa cgg aaa gag ttt gag ctg caa gta acg aca ttg aaa gat			774
Glu Gln Glu Arg Lys Glu Phe Glu Leu Gln Val Thr Thr Leu Lys Asp			
150	155	160	165
cggttcaa cat atg gaa caa cat cag aaa tca ata gtg gca tat gtt			822
Arg Leu Gln His Met Glu Gln His Gln Lys Ser Ile Val Ala Tyr Val			
170	175	180	
tca cag gtt ttg gga aaa cca gga ctt tca cta aac ctc gaa aac cat			870
Ser Gln Val Leu Gly Lys Pro Gly Leu Ser Leu Asn Leu Glu Asn His			
185	190	195	
gag aga aga aaa aga aga ttt caa gag aac tct ctt cct cca agc agt			918
Glu Arg Arg Lys Arg Arg Phe Gln Glu Asn Ser Leu Pro Pro Ser Ser			
200	205	210	
tca cac ata gaa cag gtc gaa aag tta gaa tct tcg cta acg ttt tgg			966
Ser His Ile Glu Gln Val Glu Lys Leu Glu Ser Ser Leu Thr Phe Trp			
215	220	225	
gag aat ctt gta tcg gaa tca tgc gag aag agc ggt ttg cag tca tca			1014
Glu Asn Leu Val Ser Glu Ser Cys Glu Lys Ser Gly Leu Gln Ser Ser			
230	235	240	245
agc atg gat cat gat gca gct gag tca agt cta agt att ggc gat aca			1062
Ser Met Asp His Asp Ala Ala Glu Ser Ser Leu Ser Ile Gly Asp Thr			
250	255	260	
cga ccc aaa tca tcg aag att gat atg aac tca gag cgc ccc gtt acc			1110
Arg Pro Lys Ser Ser Lys Ile Asp Met Asn Ser Glu Pro Pro Val Thr			
265	270	275	
gtt act gcg cct gct cca aaa aca ggc gtt aac gat gac ttt tgg gaa			1158
Val Thr Ala Pro Ala Pro Lys Thr Gly Val Asn Asp Asp Phe Trp Glu			
280	285	290	
caa tgt ttg aca gag aac cct gga tca acc gag caa caa gaa gtt cag			1206
Gln Cys Leu Thr Glu Asn Pro Gly Ser Thr Glu Gln Gln Glu Val Gln			
295	300	305	
tca gag aga aga gat gtc ggt aat gat aat aat ggt aat aag att gga			1254
Ser Glu Arg Arg Asp Val Gly Asn Asp Asn Asn Gly Asn Lys Ile Gly			
310	315	320	325
aat caa agg acg tat tgg tgg aat tca ggg aat gta aat aac att aca			1302

MBI15 Sequence Listing.ST25

Asn Gln Arg Thr Tyr Trp Trp Asn Ser Gly Asn Val Asn Asn Ile Thr			
330	335	340	
gag aaa gct tct tga catgaatgag gttttgtaa aatagtttc ttttggttcc 1357			
Glu Lys Ala Ser			
345			
actgagatta ttgttatgtgt tcattattta ttactctgtt tctgtaaaaa caaatctc 1417			
tattgttgaa ggcaggagtg acataaaatgc atatgcagaa ttggtttcaa aaa		1470	
<210> 40			
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Met Asp Glu Asn Asn Gly Gly Ser Ser Ser Leu Pro Pro Phe Leu Thr			
1	5	10	15
Lys Thr Tyr Glu Met Val Asp Asp Ser Ser Ser Asp Ser Val Val Ala			
20	25	30	
Trp Ser Glu Asn Asn Lys Ser Phe Ile Val Lys Asn Pro Ala Glu Phe			
35	40	45	
Ser Arg Asp Leu Leu Pro Arg Phe Phe Lys His Lys Asn Phe Ser Ser			
50	55	60	
Phe Ile Arg Gln Leu Asn Thr Tyr Gly Phe Arg Lys Val Asp Pro Glu			
65	70	75	80
Lys Trp Glu Phe Leu Asn Asp Asp Phe Val Arg Gly Arg Pro Tyr Leu			
85	90	95	
Met Lys Asn Ile His Arg Arg Lys Pro Val His Ser His Ser Leu Val			
100	105	110	
Asn Leu Gln Ala Gln Asn Pro Leu Thr Glu Ser Glu Arg Arg Ser Met			
115	120	125	
Glu Asp Gln Ile Glu Arg Leu Lys Asn Glu Lys Glu Gly Leu Leu Ala			
130	135	140	
Glu Leu Gln Asn Gln Glu Gln Glu Arg Lys Glu Phe Glu Leu Gln Val			
145	150	155	160
Thr Thr Leu Lys Asp Arg Leu Gln His Met Glu Gln His Gln Lys Ser			
165	170	175	
Ile Val Ala Tyr Val Ser Gln Val Leu Gly Lys Pro Gly Leu Ser Leu			
180	185	190	
Asn Leu Glu Asn His Glu Arg Arg Lys Arg Arg Phe Gln Glu Asn Ser			
195	200	205	
Leu Pro Pro Ser Ser Ser His Ile Glu Gln Val Glu Lys Leu Glu Ser			
210	215	220	

MBI15 Sequence Listing .ST25

Ser	Leu	Thr	Phe	Trp	Glu	Asn	Leu	Val	Ser	Glu	Ser	Cys	Glu	Lys	Ser	
225					230				235					240		
Gly Leu Gln Ser Ser Ser Met Asp His Asp Ala Ala Glu Ser Ser Leu																
					245				250					255		
Ser Ile Gly Asp Thr Arg Pro Lys Ser Ser Lys Ile Asp Met Asn Ser																
					260				265					270		
Glu Pro Pro Val Thr Val Thr Ala Pro Ala Pro Lys Thr Gly Val Asn																
					275				280					285		
Asp Asp Phe Trp Glu Gln Cys Leu Thr Glu Asn Pro Gly Ser Thr Glu																
					290				295					300		
Gln Gln Glu Val Gln Ser Glu Arg Arg Asp Val Gly Asn Asp Asn Asn																
					305				310					315		320
Gly Asn Lys Ile Gly Asn Gln Arg Thr Tyr Trp Trp Asn Ser Gly Asn																
					325				330					335		
Val Asn Asn Ile Thr Glu Lys Ala Ser																
					340				345							
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gataaatcaa tcaacaaaaac aaaaaaaaaact ctatagtttag tttctctgaa a atg tac																
														57		
														Met	Tyr	
														1		
gga cag tgc aat ata gaa tcc gac tac gct ttg ttg gag tcg ata aca																
Gly	Gln	Cys	Asn	Ile	Glu	Ser	Asp	Tyr	Ala	Leu	Leu	Glu	Ser	Ile	Thr	
5					10					15						
cgt cac ttg cta gga gga gga gag aac gag ctg cga ctc aat gag																
Arg	His	Leu	Leu	Gly	Gly	Gly	Gly	Glu	Asn	Glu	Leu	Arg	Leu	Asn	Glu	
20					25					30					153	
tca aca ccg agt tcg tgt ttc aca gag agt tgg gga ggt ttg cca ttg																
Ser	Thr	Pro	Ser	Ser	Cys	Phe	Thr	Glu	Ser	Trp	Gly	Gly	Leu	Pro	Leu	
35					40					45					201	
aaa gag aat gat tca gag gac atg ttg gtg tac gga ctc ctc aaa gat																
Lys	Glu	Asn	Asp	Ser	Glu	Asp	Met	Leu	Val	Tyr	Gly	Leu	Leu	Lys	Asp	
55					60					65					249	
gcc ttc cat ttt gac acg tca tca tcg gac ttg agc tgt ctt ttt gat																
Ala	Phe	His	Phe	Asp	Thr	Ser	Ser	Asp	Leu	Ser	Cys	Leu	Phe	Asp		
70					75					80					297	
ttt ccg gcg gtt aaa gtc gag cca act gag aac ttt acg gcg atg gag																
Phe	Pro	Ala	Val	Lys	Val	Glu	Pro	Thr	Glu	Asn	Phe	Thr	Ala	Met	Glu	
85					90					95					345	
gag aaa cca aag aaa gcg ata ccg gtt acg gag acg gca gtg aag gcg																
Glu	Lys	Pro	Lys	Lys	Ala	Ile	Pro	Val	Thr	Glu	Thr	Ala	Val	Lys	Ala	
100					105					110					393	

## MBI15 Sequence Listing.ST25

aag cat tac aga gga gtg agg cag aga ccg tgg ggg aaa ttc gcg gcg Lys His Tyr Arg Gly Val Arg Gln Arg Pro Trp Gly Lys Phe Ala Ala 115 120 125 130	441
gag ata cgt gat ccg gcg aag aat gga gct agg gtt tgg tta ggg acg Glu Ile Arg Asp Pro Ala Lys Asn Gly Ala Arg Val Trp Leu Gly Thr 135 140 145	489
ttt gag acg gcg gaa gat gcg gct tta gct tac gat ata gct gct ttt Phe Glu Thr Ala Glu Asp Ala Ala Leu Ala Tyr Asp Ile Ala Ala Phe 150 155 160	537
agg atg cgt ggt tcc cgc gct tta ttg aat ttt ccg ttg agg gtt aat Arg Met Arg Gly Ser Arg Ala Leu Leu Asn Phe Pro Leu Arg Val Asn 165 170 175	585
tcc ggt gaa cct gac ccg gtt cg <sup>g</sup> atc acg tct aag aga tct tct tcg Ser Gly Glu Pro Asp Pro Val Arg Ile Thr Ser Lys Arg Ser Ser Ser 180 185 190	633
tcg tcg tcg tcg tcc tct tct acg tcg tcg tct gaa aac aac ggg aag Ser Ser Ser Ser Ser Ser Ser Ser Ser Glu Asn Gly Lys 195 200 205 210	681
ttg aaa cga agg aga aaa gca gag aat ctg acg tcg gag gtg gtg cag Leu Lys Arg Arg Arg Lys Ala Glu Asn Leu Thr Ser Glu Val Val Gln 215 220 225	729
gtg aag tgt gag gtt ggt gat gag aca cgt gtt gat gag tta ttg gtt Val Lys Cys Glu Val Gly Asp Glu Thr Arg Val Asp Glu Leu Leu Val 230 235 240	777
tca taa gtttgatctt gtgtgttttg tagttgaata gttttgctat aaatgttgag Ser	833
gcaccaagta aaagtgttcc cgtgatgtaa attagttact aaacagagcc atatatcttc aatcaaaaaa aaaaaaaaaa	893
	913
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Met Tyr Gly Gln Cys Asn Ile Glu Ser Asp Tyr Ala Leu Leu Glu Ser 1 5 10 15	
Ile Thr Arg His Leu Leu Gly Gly Gly Glu Asn Glu Leu Arg Leu 20 25 30	
Asn Glu Ser Thr Pro Ser Ser Cys Phe Thr Glu Ser Trp Gly Gly Leu 35 40 45	
Pro Leu Lys Glu Asn Asp Ser Glu Asp Met Leu Val Tyr Gly Leu Leu 50 55 60	
Lys Asp Ala Phe His Phe Asp Thr Ser Ser Asp Leu Ser Cys Leu 65 70 75 80	
Phe Asp Phe Pro Ala Val Lys Val Glu Pro Thr Glu Asn Phe Thr Ala 85 90 95	
Met Glu Glu Lys Pro Lys Lys Ala Ile Pro Val Thr Glu Thr Ala Val 100 105 110	

## MBI15 Sequence Listing.ST25

Lys Ala Lys His Tyr Arg Gly Val Arg Gln Arg Pro Trp Gly Lys Phe  
115 120 125

Ala Ala Glu Ile Arg Asp Pro Ala Lys Asn Gly Ala Arg Val Trp Leu  
130 135 140

Gly Thr Phe Glu Thr Ala Glu Asp Ala Ala Leu Ala Tyr Asp Ile Ala  
145 150 155 160

Ala Phe Arg Met Arg Gly Ser Arg Ala Leu Leu Asn Phe Pro Leu Arg  
165 170 175

Val Asn Ser Gly Glu Pro Asp Pro Val Arg Ile Thr Ser Lys Arg Ser  
180 185 190

Ser Ser Ser Ser Ser Ser Ser Ser Thr Ser Ser Ser Glu Asn  
195 200 205

Gly Lys Leu Lys Arg Arg Lys Ala Glu Asn Leu Thr Ser Glu Val  
210 215 220

Val Gln Val Lys Cys Glu Val Gly Asp Glu Thr Arg Val Asp Glu Leu  
225 230 235 240

Leu Val Ser

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Met Thr  
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aaa tct gga gag aga cca aaa cag aga cag agg aaa ggg tta tgg tca 106  
Lys Ser Gly Glu Arg Pro Lys Gln Arg Gln Arg Lys Gly Leu Trp Ser  
5 10 15

cct gaa gaa gac cag aag ctc aag agt ttc atc ctc tct cgt ggc cat 154  
Pro Glu Glu Asp Gln Lys Leu Lys Ser Phe Ile Leu Ser Arg Gly His  
20 25 30

gct tgc tgg acc act gtt ccc atc cta gct gga ttg caa agg aat ggg 202  
Ala Cys Trp Thr Thr Val Pro Ile Leu Ala Gly Leu Gln Arg Asn Gly  
35 40 45 50

aaa agc tgc aga tta agg tgg att aat tac cta aga cca gga cta aag 250  
Lys Ser Cys Arg Leu Arg Trp Ile Asn Tyr Leu Arg Pro Gly Leu Lys  
55 60 65

agg ggg tcg ttt agt gaa gaa gaa gag acc atc ttg act tta cat 298  
Arg Gly Ser Phe Ser Glu Glu Glu Glu Thr Ile Leu Thr Leu His  
70 75 80

tct tcc ttg ggt aac aag tgg tct cgg att gca aaa tat tta ccg gga 346

MBI15 Sequence Listing.ST25

Ser Ser Leu Gly Asn Lys Trp Ser Arg Ile Ala Lys Tyr Leu Pro Gly			
85	90	95	
aga aca gac aac gag att aag aac tat tgg cat tcc tat ctg aag aag	394		
Arg Thr Asp Asn Glu Ile Lys Asn Tyr Trp His Ser Tyr Leu Lys Lys			
100	105	110	
aga tgg ctc aaa tct caa cca caa ctc aaa agc caa ata tca gac ctc	442		
Arg Trp Leu Lys Ser Gln Pro Gln Leu Lys Ser Gln Ile Ser Asp Leu			
115	120	125	130
aca gaa tct cct tct tca cta ctt tct tgc ggg aaa aga aat ctg gaa	490		
Thr Glu Ser Pro Ser Ser Leu Leu Ser Cys Gly Lys Arg Asn Leu Glu			
135	140	145	
acc gaa acc cta gat cac gtg atc tcc ttc cag aaa ttt tca gag aat	538		
Thr Glu Thr Leu Asp His Val Ile Ser Phe Gln Lys Phe Ser Glu Asn			
150	155	160	
cca act tca tca cca tcc aaa gaa agc aac aac aac atg atc atg aac	586		
Pro Thr Ser Ser Pro Ser Lys Glu Ser Asn Asn Asn Met Ile Met Asn			
165	170	175	
aac agt aat aac ttg cct aaa ctg ttc ttc tct gag tgg atc agt tct	634		
Asn Ser Asn Asn Leu Pro Lys Leu Phe Phe Ser Glu Trp Ile Ser Ser			
180	185	190	
tca aat cca cac atc gat tac tcc tct gct ttt aca gat tcc aag cac	682		
Ser Asn Pro His Ile Asp Tyr Ser Ser Ala Phe Thr Asp Ser Lys His			
195	200	205	210
att aat gaa actcaa gat caa atc aat gaa gag gaa gtg atg atg atc	730		
Ile Asn Glu Thr Gln Asp Gln Ile Asn Glu Glu Glu Val Met Met Ile			
215	220	225	
aat aac aac tac tct tca ctt gag gat gtc atg ctc cgt aca gat	778		
Asn Asn Asn Tyr Ser Ser Leu Glu Asp Val Met Leu Arg Thr Asp			
230	235	240	
ttt ttg cag cct gat cat gaa tat gca aat tat tat tct tct gga gat	826		
Phe Leu Gln Pro Asp His Glu Tyr Ala Asn Tyr Tyr Ser Ser Gly Asp			
245	250	255	
ttc ttc atc aac agt gac caa aat tat gtc taa gaagagtcaa tatgatcgta	879		
Phe Ile Asn Ser Asp Gln Asn Tyr Val			
260	265		
agaggaacat aagcttagtta cttgtgttac agc	912		
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Met Thr Lys Ser Gly Glu Arg Pro Lys Gln Arg Gln Lys Gly Leu			
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Trp Ser Pro Glu Glu Asp Gln Lys Leu Lys Ser Phe Ile Leu Ser Arg			
20 25 30			
Gly His Ala Cys Trp Thr Val Pro Ile Leu Ala Gly Leu Gln Arg			
35 40 45			
Asn Gly Lys Ser Cys Arg Leu Arg Trp Ile Asn Tyr Leu Arg Pro Gly			
50 55 60			
Leu Lys Arg Gly Ser Phe Ser Glu Glu Glu Glu Thr Ile Leu Thr			
65 70 75 80			

## MBI15 Sequence Listing.ST25

Leu His Ser Ser Leu Gly Asn Lys Trp Ser Arg Ile Ala Lys Tyr Leu  
85 90 95

Pro Gly Arg Thr Asp Asn Glu Ile Lys Asn Tyr Trp His Ser Tyr Leu  
100 105 110

Lys Lys Arg Trp Leu Lys Ser Gln Pro Gln Leu Lys Ser Gln Ile Ser  
115 120 125

Asp Leu Thr Glu Ser Pro Ser Ser Leu Leu Ser Cys Gly Lys Arg Asn  
130 135 140

Leu Glu Thr Glu Thr Leu Asp His Val Ile Ser Phe Gln Lys Phe Ser  
145 150 155 160

Glu Asn Pro Thr Ser Ser Pro Ser Lys Glu Ser Asn Asn Asn Met Ile  
165 170 175

Met Asn Asn Ser Asn Asn Leu Pro Lys Leu Phe Phe Ser Glu Trp Ile  
180 185 190

Ser Ser Ser Asn Pro His Ile Asp Tyr Ser Ser Ala Phe Thr Asp Ser  
195 200 205

Lys His Ile Asn Glu Thr Gln Asp Gln Ile Asn Glu Glu Glu Val Met  
210 215 220

Met Ile Asn Asn Asn Asn Tyr Ser Ser Leu Glu Asp Val Met Leu Arg  
225 230 235 240

Thr Asp Phe Leu Gln Pro Asp His Glu Tyr Ala Asn Tyr Tyr Ser Ser  
245 250 255

Gly Asp Phe Phe Ile Asn Ser Asp Gln Asn Tyr Val  
260 265

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<223> G2550

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Met Ala Val Tyr Tyr Pro Asn Ser Val Gly Met Gln Ser Leu Tyr Gln  
1 5 10 15

gaa tcc att tac ctc aac gaa caa caa caa caa caa caa gct tct 96  
Glu Ser Ile Tyr Leu Asn Glu Gln Gln Gln Gln Gln Ala Ser  
20 25 30

tct tcc tct gct gca tct ttc tcc gag att gtt tcc ggt gat gtt cga 144  
Ser Ser Ser Ala Ala Ser Phe Ser Glu Ile Val Ser Gly Asp Val Arg  
35 40 45

aac aac gag atg gta ttt atc cca cca aca agc gac gta gcc gtc aac 192

MBI15 Sequence Listing, ST25

Asn Asn Glu Met Val Phe Ile Pro Pro Thr Ser Asp Val Ala Val Asn				
50	55	60		
gga aac gta acg gtg tca agt aac gat cta agc ttt cac ggt gga gga				240
Gly Asn Val Thr Val Ser Ser Asn Asp Leu Ser Phe His Gly Gly				
65	70	75	80	
ctt tct tta agt ctt ggt aat cag atc cag tca gct gtc tct gtt tct				288
Leu Ser Leu Ser Leu Gly Asn Gln Ile Gln Ser Ala Val Ser Val Ser				
85	90	95		
ccg ttt cag tat cat tac cag aat ctt tcg aac caa ttg agt tac aat				336
Pro Phe Gln Tyr His Tyr Gln Asn Leu Ser Asn Gln Leu Ser Tyr Asn				
100	105	110		
aat ctt aat cct tct act atg tct gat gag aat ggg aag agc ttg agt				384
Asn Leu Asn Pro Ser Thr Met Ser Asp Glu Asn Gly Lys Ser Leu Ser				
115	120	125		
gtt cat cag cat cac tct gat caa att tta cct tcc tct gtt tac aac				432
Val His Gln His His Ser Asp Gln Ile Leu Pro Ser Ser Val Val Tyr Asn				
130	135	140		
aac aat ggt aat aat ggt gtt gga ttc tac aac aat tac cgt tac gag				480
Asn Asn Gly Asn Asn Gly Val Gly Phe Tyr Asn Asn Tyr Arg Tyr Glu				
145	150	155	160	
aca tca ggg ttt gtg agt agt gta ctg aga tct cgt tac ctt aaa cca				528
Thr Ser Gly Phe Val Ser Ser Val Leu Arg Ser Arg Tyr Leu Lys Pro				
165	170	175		
aca caa caa ttg ctt gat gaa gtt gtt agt gta agg aaa gat ttg aaa				576
Thr Gln Gln Leu Leu Asp Glu Val Val Ser Val Arg Lys Asp Leu Lys				
180	185	190		
ttg ggg aat aag aag atg aag aat gat aaa ggt caa gac ttt cac aat				624
Leu Gly Asn Lys Lys Met Lys Asn Asp Lys Gly Gln Asp Phe His Asn				
195	200	205		
ggg tct agt gat aac att aca gaa gat gat aaa tct caa tcg cag gag				672
Gly Ser Ser Asp Asn Ile Thr Glu Asp Asp Lys Ser Gln Ser Gln Glu				
210	215	220		
ttg tct cct tca gaa cgt cag gag cta cag agc aag aag agc aag ctt				720
Leu Ser Pro Ser Glu Arg Gln Glu Leu Gln Ser Lys Lys Ser Lys Leu				
225	230	235	240	
tta aca atg gtg gat gag gta gat aaa agg tat aac caa tac cat cat				768
Leu Thr Met Val Asp Glu Val Asp Lys Arg Tyr Asn Gln Tyr His His				
245	250	255		
caa atg gaa gct tta gca tcg tct ttc gag atg gta aca ggt ctt gga				816
Gln Met Glu Ala Leu Ala Ser Ser Phe Glu Met Val Thr Gly Leu Gly				
260	265	270		
gca gct aag cct tac aca tcc gta gct ctg aat aga atc tct cgc cat				864
Ala Ala Lys Pro Tyr Thr Ser Val Ala Leu Asn Arg Ile Ser Arg His				
275	280	285		
ttc cgc tgt tta cgc gac gcg ata aaa gaa cag att cag gtg atc aga				912
Phe Arg Cys Leu Arg Asp Ala Ile Lys Glu Gln Ile Gln Val Ile Arg				
290	295	300		
ggg aag ctt ggg gag aga gag act tct gat gaa caa gga gag agg ata				960
Gly Lys Leu Gly Glu Arg Glu Thr Ser Asp Glu Gln Gly Glu Arg Ile				
305	310	315	320	
ccg cgt ctt agg tac tta gat caa cgg ttg aga caa cag aga gct ttg				1008
Pro Arg Leu Arg Tyr Leu Asp Gln Arg Leu Arg Gln Gln Arg Ala Leu				
325	330	335		
cat caa caa ctt gga atg gtt aga cca gct tgg aga cca caa aga ggc				1056
His Gln Gln Leu Gly Met Val Arg Pro Ala Trp Arg Pro Gln Arg Gly				
340	345	350		

MBI15 Sequence Listing.ST25

tta cct gaa aac tct gtc tct ata ctt cga gct tgg ctc ttt gag cat Leu Pro Glu Asn Ser Val Ser Ile Leu Arg Ala Trp Leu Phe Glu His 355 360 365	1104
ttc ctt cat cca tat cct aaa gaa tca gag aaa atc atg ctt tca aag Phe Leu His Pro Tyr Pro Lys Glu Ser Glu Lys Ile Met Leu Ser Lys 370 375 380	1152
cag aca gga cta tcg aaa aac cag gtt gca aat tgg ttt att aac gcg Gln Thr Gly Leu Ser Lys Asn Gln Val Ala Asn Trp Phe Ile Asn Ala 385 390 395 400	1200
aga gtt cga cta tgg aaa cca atg att gaa gag atg tat aaa gaa gag Arg Val Arg Leu Trp Lys Pro Met Ile Glu Met Tyr Lys Glu Glu 405 410 415	1248
ttt gga gaa tca gca gag tta ctc tct aac tct aat caa gac acc aaa Phe Gly Glu Ser Ala Glu Leu Leu Ser Asn Ser Asn Gln Asp Thr Lys 420 425 430	1296
aaa atg cag gaa aca tct cag ctc aaa cac gaa gac tct tcg tct tcg Lys Met Gln Glu Thr Ser Gln Leu Lys His Glu Asp Ser Ser Ser Ser 435 440 445	1344
caa caa cag aat cag gga aac aac aac aac atc cca tat aca tct Gln Gln Gln Asn Gln Gly Asn Asn Asn Asn Ile Pro Tyr Thr Ser 450 455 460	1392
gat gca gaa caa aac cta gtc ttt gca gat cct aaa cca gac cgt gct Asp Ala Glu Gln Asn Leu Val Phe Ala Asp Pro Lys Pro Asp Arg Ala 465 470 475 480	1440
act act gga gat tac gac agc ttg atg aac tat cat ggg ttt ggt att Thr Thr Gly Asp Tyr Asp Ser Leu Met Asn Tyr His Gly Phe Gly Ile 485 490 495	1488
gat gat tac aat cgt tac gtt ggc ctt gga aac caa caa gat ggc aga Asp Asp Tyr Asn Arg Tyr Val Gly Leu Gly Asn Gln Gln Asp Gly Arg 500 505 510	1536
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Glu Ser Ile Tyr Leu Asn Glu Gln Gln Gln Gln Gln Ala Ser 20 25 30	
Ser Ser Ser Ala Ala Ser Phe Ser Glu Ile Val Ser Gly Asp Val Arg 35 40 45	
Asn Asn Glu Met Val Phe Ile Pro Pro Thr Ser Asp Val Ala Val Asn 50 55 60	
Gly Asn Val Thr Val Ser Ser Asn Asp Leu Ser Phe His Gly Gly Gly 65 70 75 80	
Leu Ser Leu Ser Leu Gly Asn Gln Ile Gln Ser Ala Val Ser Val Ser 85 90 95	

## MBI15 Sequence Listing .ST25

Pro Phe Gln Tyr His Tyr Gln Asn Leu Ser Asn Gln Leu Ser Tyr Asn  
100 105 110

Asn Leu Asn Pro Ser Thr Met Ser Asp Glu Asn Gly Lys Ser Leu Ser  
115 120 125

Val His Gln His His Ser Asp Gln Ile Leu Pro Ser Ser Val Tyr Asn  
130 135 140

Asn Asn Gly Asn Asn Gly Val Gly Phe Tyr Asn Asn Tyr Arg Tyr Glu  
145 150 155 160

Thr Ser Gly Phe Val Ser Ser Val Leu Arg Ser Arg Tyr Leu Lys Pro  
165 170 175

Thr Gln Gln Leu Leu Asp Glu Val Val Ser Val Arg Lys Asp Leu Lys  
180 185 190

Leu Gly Asn Lys Lys Met Lys Asn Asp Lys Gly Gln Asp Phe His Asn  
195 200 205

Gly Ser Ser Asp Asn Ile Thr Glu Asp Asp Lys Ser Gln Ser Gln Glu  
210 215 220

Leu Ser Pro Ser Glu Arg Gln Glu Leu Gln Ser Lys Lys Ser Lys Leu  
225 230 235 240

Leu Thr Met Val Asp Glu Val Asp Lys Arg Tyr Asn Gln Tyr His His  
245 250 255

Gln Met Glu Ala Leu Ala Ser Ser Phe Glu Met Val Thr Gly Leu Gly  
260 265 270

Ala Ala Lys Pro Tyr Thr Ser Val Ala Leu Asn Arg Ile Ser Arg His  
275 280 285

Phe Arg Cys Leu Arg Asp Ala Ile Lys Glu Gln Ile Gln Val Ile Arg  
290 295 300

Gly Lys Leu Gly Glu Arg Glu Thr Ser Asp Glu Gln Gly Glu Arg Ile  
305 310 315 320

Pro Arg Leu Arg Tyr Leu Asp Gln Arg Leu Arg Gln Gln Arg Ala Leu  
325 330 335

His Gln Gln Leu Gly Met Val Arg Pro Ala Trp Arg Pro Gln Arg Gly  
340 345 350

Leu Pro Glu Asn Ser Val Ser Ile Leu Arg Ala Trp Leu Phe Glu His  
355 360 365

Phe Leu His Pro Tyr Pro Lys Glu Ser Glu Lys Ile Met Leu Ser Lys  
370 375 380

Gln Thr Gly Leu Ser Lys Asn Gln Val Ala Asn Trp Phe Ile Asn Ala  
385 390 395 400

## MBI15 Sequence Listing.ST25

Arg Val Arg Leu Trp Lys Pro Met Ile Glu Glu Met Tyr Lys Glu Glu  
 405 410 415

Phe Gly Glu Ser Ala Glu Leu Leu Ser Asn Ser Asn Gln Asp Thr Lys  
 420 425 430

Lys Met Gln Glu Thr Ser Gln Leu Lys His Glu Asp Ser Ser Ser Ser  
 435 440 445

Gln Gln Gln Asn Gln Gly Asn Asn Asn Asn Ile Pro Tyr Thr Ser  
 450 455 460

Asp Ala Glu Gln Asn Leu Val Phe Ala Asp Pro Lys Pro Asp Arg Ala  
 465 470 475 480

Thr Thr Gly Asp Tyr Asp Ser Leu Met Asn Tyr His Gly Phe Gly Ile  
 485 490 495

Asp Asp Tyr Asn Arg Tyr Val Gly Leu Gly Asn Gln Gln Asp Gly Arg  
 500 505 510

Tyr Ser Asn Pro His Gln Leu His Asp Phe Val Val  
 515 520

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<223> G965

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Met Gly Leu Ala Thr Thr Ser Ser Met Ser Gln Asp  
1 5 10

tat cat cat cac caa gga atc ttt tcc ttc tct aat gga ttc cac cga 159  
Tyr His His His Gln Gly Ile Phe Ser Phe Ser Asn Gly Phe His Arg  
15 20 25

tca tca tca acc act cat cag gag gaa gta gat gaa tcc gcc gtc gtc 207  
Ser Ser Ser Thr Thr His Gln Glu Glu Val Asp Glu Ser Ala Val Val  
30 35 40 45

tcc ggt gct caa att ccg gtt tat gaa acc gcc gga atg ttg tct gaa 255  
Ser Gly Ala Gln Ile Pro Val Tyr Glu Thr Ala Gly Met Leu Ser Glu  
50 55 60

atg ttt gct tac cct ggc gga ggt ggc ggc ggt tcc ggt gga gag att 303  
Met Phe Ala Tyr Pro Gly Gly Gly Gly Ser Gly Gly Glu Ile  
65 70 75

ctt gat cag tct act aaa cag ttg cta gag caa caa aac cgt cac aac 351  
Leu Asp Gln Ser Thr Lys Gln Leu Leu Glu Gln Gln Asn Arg His Asn  
80 85 90

aac aac aat aac tca act ctt cat atg tta tta cca aat cat cat caa 399  
Asn Asn Asn Ser Thr Leu His Met Leu Leu Pro Asn His His Gln  
95 100 105

## MBI15 Sequence Listing.ST25

gg ttt gct ttc acc gac gaa aac act atg cag ccg cag caa caa caa Gly Phe Ala Phe Thr Asp Glu Asn Thr Met Gln Pro Gln Gln Gln 110 115 120 125	447
cac ttt aca tgg cca tct tcc tcc gat cat cat caa aac cga gat His Phe Thr Trp Pro Ser Ser Ser Asp His His Gln Asn Arg Asp 130 135 140	495
atg atc gga acc gtc cac gtg gaa gga aag ggt ttg tct tta tct Met Ile Gly Thr Val His Val Glu Gly Lys Gly Leu Ser Leu Ser 145 150 155	543
ctc tca tct tca tta gcc gca gct aaa gcc gag gaa tat aga agc att Leu Ser Ser Ser Leu Ala Ala Lys Ala Glu Glu Tyr Arg Ser Ile 160 165 170	591
tat tgt gca gcc gtt gat gga act tct tct tct aac gca tcc gct Tyr Cys Ala Ala Val Asp Gly Thr Ser Ser Ser Asn Ala Ser Ala 175 180 185	639
cat cat cat caa ttc aat cag ttc aag aat ctt ctt ctt gag aat tct His His His Gln Phe Asn Gln Phe Lys Asn Leu Leu Glu Asn Ser 190 195 200 205	687
tct tct caa cat cat cac cat caa gtt gtt gga cat ttt ggt tca tca Ser Ser Gln His His Gln Val Val Gly His Phe Gly Ser Ser 210 215 220	735
tca tca tct ccc atg gcg gct tct tca tcc att gga ggg atc tac acg Ser Ser Ser Pro Met Ala Ala Ser Ser Ser Ile Gly Gly Ile Tyr Thr 225 230 235	783
ttg agg aat tcg aaa tat acg aaa ccg gct caa gag ttg ttg gaa gag Leu Arg Asn Ser Lys Tyr Thr Lys Pro Ala Gln Glu Leu Leu Glu Glu 240 245 250	831
ttt tgt agt gtt gga aga gga cat ttc aag aag aac aaa ctt agt agg Phe Cys Ser Val Gly Arg Gly His Phe Lys Lys Asn Lys Leu Ser Arg 255 260 265	879
aac aac tca aac cct aat act acc ggt gga gga ggc gga ggg tcc Asn Asn Ser Asn Pro Asn Thr Thr Gly Gly Gly Gly Gly Ser 270 275 280 285	927
tcg tca tcg gcc gga aca gct aat gat agt cct cct ttg tct ccg gct Ser Ser Ser Ala Gly Thr Ala Asn Asp Ser Pro Pro Leu Ser Pro Ala 290 295 300	975
gat cgg att gaa cat caa aga aga aaa gtc aag cta cta tct atg ctt Asp Arg Ile Glu His Gln Arg Arg Lys Val Lys Leu Leu Ser Met Leu 305 310 315	1023
gaa gag gtg gac cga cgg tac aac cac tac tgc gaa caa atg caa atg Glu Glu Val Asp Arg Arg Tyr Asn His Tyr Cys Glu Gln Met Gln Met 320 325 330	1071
gta gtg aac tca ttc gac caa gta atg ggt tac ggc gcg gcg gtt ccg Val Val Asn Ser Phe Asp Gln Val Met Gly Tyr Gly Ala Ala Val Pro 335 340 345	1119
tac acg aca tta gct caa aag gca atg tct agg cat ttc ccg tgt ttg Tyr Thr Thr Leu Ala Gln Lys Ala Met Ser Arg His Phe Arg Cys Leu 350 355 360 365	1167
aaa gac gcg gta gcg gtt cag ctt aaa cgc agc tgt gag ctt cta ggg Lys Asp Ala Val Ala Val Gln Leu Lys Arg Ser Cys Glu Leu Leu Gly 370 375 380	1215
gat aaa gag gcg gca ggg gct gca tcc tcg ggg tta acc aaa ggg gaa Asp Lys Glu Ala Ala Gly Ala Ala Ser Ser Gly Leu Thr Lys Gly Glu 385 390 395	1263
acg ccg cga ttg cgt ttg cta gag cag agt ttg cgt cag caa cga gcg Thr Pro Arg Leu Arg Leu Leu Glu Gln Ser Leu Arg Gln Gln Arg Ala	1311

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400

405

410

ttt cat cat atg ggt atg atg gag caa gag gca tgg aga ccg caa cgt      1359  
 Phe His His Met Gly Met Met Glu Gln Glu Ala Trp Arg Pro Gln Arg  
 415                          420                          425

ggt ttg cct gaa cgc tcc gtt aat atc ctt aga gct tgg cta ttc gag      1407  
 Gly Leu Pro Glu Arg Ser Val Asn Ile Leu Arg Ala Trp Leu Phe Glu  
 430                          435                          440                          445

cat ttt ctt aat ccg tac cca agc gat gct gat aag cac ctc tta gca      1455  
 His Phe Leu Asn Pro Tyr Pro Ser Asp Ala Asp Lys His Leu Leu Ala  
 450                          455                          460

cga cag act ggt tta tcc aga aat cag gtg tca aat tgg ttc ata aat      1503  
 Arg Gln Thr Gly Leu Ser Arg Asn Gln Val Ser Asn Trp Phe Ile Asn  
 465                          470                          475

gct agg gtt cgc cta tgg aaa cca atg gtg gaa gag atg tat caa caa      1551  
 Ala Arg Val Arg Leu Trp Lys Pro Met Val Glu Glu Met Tyr Gln Gln  
 480                          485                          490

gaa gca aaa gaa aga gaa gca gaa gaa aat gaa aat caa caa      1599  
 Glu Ala Lys Glu Arg Glu Ala Glu Glu Glu Asn Glu Asn Gln Gln  
 495                          500                          505

caa caa aga aga cag caa caa aca aac aac gac acg aaa ccc aac      1647  
 Gln Gln Arg Arg Gln Gln Gln Thr Asn Asn Asn Asp Thr Lys Pro Asn  
 510                          515                          520                          525

aac aat gaa aac aac ttc act gtc ata acc gca caa act cca acg acg      1695  
 Asn Asn Glu Asn Asn Phe Thr Val Ile Thr Ala Gln Thr Pro Thr Thr  
 530                          535                          540

atg aca tcg aca cat cac gaa aac gac tct tca ttc ctc tct tcc gtc      1743  
 Met Thr Ser Thr His His Glu Asn Asp Ser Ser Phe Leu Ser Ser Val  
 545                          550                          555

gcc gcc gct tct cac ggc ggt tca gac gcg ttc acc gtc gcc acg tgt      1791  
 Ala Ala Ser His Gly Gly Ser Asp Ala Phe Thr Val Ala Thr Cys  
 560                          565                          570

cag caa gac gtc agt gac ttc cac gtc gac gga gat ggt gtg aac gtc      1839  
 Gln Gln Asp Val Ser Asp Phe His Val Asp Gly Asp Gly Val Asn Val  
 575                          580                          585

ata aga ttc ggg acc aaa cag act ggt gac gtg tct ctt acg ctt ggt      1887  
 Ile Arg Phe Gly Thr Lys Gln Thr Gly Asp Val Ser Leu Thr Leu Gly  
 590                          595                          600                          605

cta cgc cac tct ggc aat att cct gat aag aac act tct ttc tcc gtt      1935  
 Leu Arg His Ser Gly Asn Ile Pro Asp Lys Asn Thr Ser Phe Ser Val  
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aga gac ttt gga gat ttt tag tcttctttgt ttctcaattt attcatc      1983  
 Arg Asp Phe Gly Asp Phe  
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<400> 48

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Thr Thr His Gln Glu Glu Val Asp Glu Ser Ala Val Val Ser Gly Ala

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35                          40                          45

Gln Ile Pro Val Tyr Glu Thr Ala Gly Met Leu Ser Glu Met Phe Ala  
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Tyr Pro Gly Gly Gly Gly Ser Gly Gly Glu Ile Leu Asp Gln  
65                          70                          75                          80

Ser Thr Lys Gln Leu Leu Glu Gln Gln Asn Arg His Asn Asn Asn  
85                          90                          95

Asn Ser Thr Leu His Met Leu Leu Pro Asn His His Gln Gly Phe Ala  
100                          105                          110

Phe Thr Asp Glu Asn Thr Met Gln Pro Gln Gln Gln His Phe Thr  
115                          120                          125

Trp Pro Ser Ser Ser Asp His His Gln Asn Arg Asp Met Ile Gly  
130                          135                          140

Thr Val His Val Glu Gly Gly Lys Gly Leu Ser Leu Ser Leu Ser Ser  
145                          150                          155                          160

Ser Leu Ala Ala Ala Lys Ala Glu Glu Tyr Arg Ser Ile Tyr Cys Ala  
165                          170                          175

Ala Val Asp Gly Thr Ser Ser Ser Asn Ala Ser Ala His His His  
180                          185                          190

Gln Phe Asn Gln Phe Lys Asn Leu Leu Glu Asn Ser Ser Ser Gln  
195                          200                          205

His His His His Gln Val Val Gly His Phe Gly Ser Ser Ser Ser  
210                          215                          220

Pro Met Ala Ala Ser Ser Ser Ile Gly Gly Ile Tyr Thr Leu Arg Asn  
225                          230                          235                          240

Ser Lys Tyr Thr Lys Pro Ala Gln Glu Leu Leu Glu Glu Phe Cys Ser  
245                          250                          255

Val Gly Arg Gly His Phe Lys Lys Asn Lys Leu Ser Arg Asn Asn Ser  
260                          265                          270

Asn Pro Asn Thr Thr Gly Gly Gly Gly Ser Ser Ser Ser  
275                          280                          285

Ala Gly Thr Ala Asn Asp Ser Pro Pro Leu Ser Pro Ala Asp Arg Ile  
290                          295                          300

Glu His Gln Arg Arg Lys Val Lys Leu Leu Ser Met Leu Glu Glu Val  
305                          310                          315                          320

Asp Arg Arg Tyr Asn His Tyr Cys Glu Gln Met Gln Met Val Val Asn  
325                          330                          335

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Ser Phe Asp Gln Val Met Gly Tyr Gly Ala Ala Val Pro Tyr Thr Thr  
340 345 350

Leu Ala Gln Lys Ala Met Ser Arg His Phe Arg Cys Leu Lys Asp Ala  
355 360 365

Val Ala Val Gln Leu Lys Arg Ser Cys Glu Leu Leu Gly Asp Lys Glu  
370 375 380

Ala Ala Gly Ala Ala Ser Ser Gly Leu Thr Lys Gly Glu Thr Pro Arg  
385 390 395 400

Leu Arg Leu Leu Glu Gln Ser Leu Arg Gln Gln Arg Ala Phe His His  
405 410 415

Met Gly Met Met Glu Gln Glu Ala Trp Arg Pro Gln Arg Gly Leu Pro  
420 425 430

Glu Arg Ser Val Asn Ile Leu Arg Ala Trp Leu Phe Glu His Phe Leu  
435 440 445

Asn Pro Tyr Pro Ser Asp Ala Asp Lys His Leu Leu Ala Arg Gln Thr  
450 455 460

Gly Leu Ser Arg Asn Gln Val Ser Asn Trp Phe Ile Asn Ala Arg Val  
465 470 475 480

Arg Leu Trp Lys Pro Met Val Glu Met Tyr Gln Gln Glu Ala Lys  
485 490 495

Glu Arg Glu Glu Ala Glu Glu Asn Glu Asn Gln Gln Gln Arg  
500 505 510

Arg Gln Gln Gln Thr Asn Asn Asp Thr Lys Pro Asn Asn Asn Glu  
515 520 525

Asn Asn Phe Thr Val Ile Thr Ala Gln Thr Pro Thr Thr Met Thr Ser  
530 535 540

Thr His His Glu Asn Asp Ser Ser Phe Leu Ser Ser Val Ala Ala Ala  
545 550 555 560

Ser His Gly Gly Ser Asp Ala Phe Thr Val Ala Thr Cys Gln Gln Asp  
565 570 575

Val Ser Asp Phe His Val Asp Gly Asp Gly Val Asn Val Ile Arg Phe  
580 585 590

Gly Thr Lys Gln Thr Gly Asp Val Ser Leu Thr Leu Gly Leu Arg His  
595 600 605

Ser Gly Asn Ile Pro Asp Lys Asn Thr Ser Phe Ser Val Arg Asp Phe  
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Gly Asp Phe  
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## MBI15 Sequence Listing .ST25

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<223> G793

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ctttactcgt ttccttc atg gct aat aac aac atc cca cat gat agc      170
    Met Ala Asn Asn Asn Ile Pro His Asp Ser
    1           5           10

atc tcc gat cca tct cct acc gac gat ttc ttc gag cag atc ctc ggg      218
Ile Ser Asp Pro Ser Pro Thr Asp Asp Phe Phe Glu Gln Ile Leu Gly
    15          20          25

ctt tcc aac ttc tcc ggt tct tca ggt tct ggt ctc tct gga atc ggc      266
Leu Ser Asn Phe Ser Gly Ser Ser Gly Ser Gly Leu Ser Gly Ile Gly
    30          35          40

ggc gtg ggt cca cct ccg atg atg ctt cag ctt ggt tca ggc aac gaa      314
Gly Val Gly Pro Pro Met Met Leu Gln Leu Gly Ser Gly Asn Glu
    45          50          55

ggg aat cat aat cat atg ggt gcc att gga gga ggt gga cct gta ggg      362
Gly Asn His Asn His Met Gly Ala Ile Gly Gly Gly Pro Val Gly
    60          65          70          75

ttt cat aat cag atg ttt ccg ttg gga tta agt ctc gat caa ggg aaa      410
Phe His Asn Gln Met Phe Pro Leu Gly Leu Ser Leu Asp Gln Gly Lys
    80          85          90

gga cat ggc ttt ctt aaa cct gat gaa act ggt aaa cgt ttc caa gac      458
Gly His Gly Phe Leu Lys Pro Asp Glu Thr Gly Lys Arg Phe Gln Asp
    95          100         105

gat gtt ctt gat aat cga tgt tcc tct atg aaa cct att ttc cat ggg      506
Asp Val Leu Asp Asn Arg Cys Ser Ser Met Lys Pro Ile Phe His Gly
    110         115         120

cag cca atg tca cag cca gct cca cca atg ccg cat caa cag tct act      554
Gln Pro Met Ser Gln Pro Ala Pro Pro Met Pro His Gln Gln Ser Thr
    125         130         135

att cggtt cct aga gtt agg gct agg cga ggt caa gct acc gat cca cat      602
Ile Arg Pro Arg Val Arg Ala Arg Arg Gly Gln Ala Thr Asp Pro His
    140         145         150         155

agc atc gct gag agg ctc cga agg gaa aga ata gca gaa cgg atc agg      650
Ser Ile Ala Glu Arg Leu Arg Arg Glu Arg Ile Ala Glu Arg Ile Arg
    160         165         170

tcg ttg cag gaa ctt gta cct acc gtt aac aag aca gat agg gct gct      698
Ser Leu Gln Glu Leu Val Pro Thr Val Asn Lys Thr Asp Arg Ala Ala
    175         180         185

atg atc gac gag att gtc gat tat gta aag ttt ctc agg ctc caa gtt      746
Met Ile Asp Glu Ile Val Asp Tyr Val Lys Phe Leu Arg Leu Gln Val
    190         195         200

aag gtc ctg agc atg agc cgt ctt ggt gga gcc ggt gct gtc gca cca      794
Lys Val Leu Ser Met Ser Arg Leu Gly Gly Ala Gly Ala Val Ala Pro
    205         210         215

cta gtc act gaa atg cca tta tct tca tca gtt gag gat gag acg cag      842
Leu Val Thr Glu Met Pro Leu Ser Ser Val Glu Asp Glu Thr Gln

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## MBI15 Sequence Listing .ST25

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240	245	250														
aag	ctg	atg	gaa	gaa	aac	gtt	gga	gca	gcg	atg	caa	ctt	ttg	caa	tca	938
Lys	Leu	Met	Glu	Glu	Asn	Val	Gly	Ala	Ala	Met	Gln	Leu	Leu	Gln	Ser	
255	260	265														
aag	gct	ctt	tgc	ata	atg	ccg	atc	tca	ttg	gca	atg	gcg	att	tac	cat	986
Lys	Ala	Leu	Cys	Ile	Met	Pro	Ile	Ser	Leu	Ala	Met	Ala	Ile	Tyr	His	
270	275	280														
tct	cag	cca	cca	gac	aca	tct	tct	tca	atc	gtc	aaa	cca	gag	atg	aat	1034
Ser	Gln	Pro	Pro	Asp	Thr	Ser	Ser	Ser	Ser	Ile	Val	Lys	Pro	Glu	Met	Asn
285	290	295														
cct	cca	ccg	tag	atttttgttc	atccaaacggt	ccccagctga	tgattgacat		1086							
Pro	Pro	Pro														
300																
tttgctctgt	ttcccaactac	tagactttt	tgactcatga	aaggtaagta	aaaaggcatt		1146									
ggagatggaa	tctaagttagg	atttgtcag	taaagaagta	aaacgggatc	tgtcaaaaga		1206									
aggaaaaaagc	tctcgcttgc	ttggctagta	tttatcattt	tgatgaaagt	aactctttt		1266									
tgttcaaaaga	ctttagtgtg	attttcagga	ccaaggcct	tgagggtagt	gctagctgta		1326									
gtaatagtaa	tgaaggtgt	ggatcgtgtt	tttgaattat	gtaaaaaagg	aagaaaaaac		1386									
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<212> PRT  
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<400> 50

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20						25					30				

Gly	Ser	Ser	Gly	Ser	Gly	Leu	Ser	Gly	Ile	Gly	Gly	Val	Gly	Pro	Pro
35						40					45				

Pro	Met	Met	Leu	Gln	Leu	Gly	Ser	Gly	Asn	Glu	Gly	Asn	His	Asn	His
50						55			60						

Met	Gly	Ala	Ile	Gly	Gly	Gly	Pro	Val	Gly	Phe	His	Asn	Gln	Met
65						70			75			80		

Phe	Pro	Leu	Gly	Leu	Ser	Leu	Asp	Gln	Gly	Lys	Gly	His	Gly	Phe	Leu
85							90					95			

Lys	Pro	Asp	Glu	Thr	Gly	Lys	Arg	Phe	Gln	Asp	Asp	Val	Leu	Asp	Asn
100							105					110			

Arg	Cys	Ser	Ser	Met	Lys	Pro	Ile	Phe	His	Gly	Gln	Pro	Met	Ser	Gln
115							120				125				

Pro Ala Pro Pro Met Pro His Gln Gln Ser Thr Ile Arg Pro Arg Val

MBI15 Sequence Listing.ST25

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Arg Ala Arg Arg Gly Gln Ala Thr Asp Pro His Ser Ile Ala Glu Arg		
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Leu Arg Arg Glu Arg Ile Ala Glu Arg Ile Arg Ser Leu Gln Glu Leu		
165	170	175
Val Pro Thr Val Asn Lys Thr Asp Arg Ala Ala Met Ile Asp Glu Ile		
180	185	190
Val Asp Tyr Val Lys Phe Leu Arg Leu Gln Val Lys Val Leu Ser Met		
195	200	205
Ser Arg Leu Gly Gly Ala Gly Ala Val Ala Pro Leu Val Thr Glu Met		
210	215	220
Pro Leu Ser Ser Ser Val Glu Asp Glu Thr Gln Ala Val Trp Glu Lys		
225	230	235
Trp Ser Asn Asp Gly Thr Glu Arg Gln Val Ala Lys Leu Met Glu Glu		
245	250	255
Asn Val Gly Ala Ala Met Gln Leu Leu Gln Ser Lys Ala Leu Cys Ile		
260	265	270
Met Pro Ile Ser Leu Ala Met Ala Ile Tyr His Ser Gln Pro Pro Asp		
275	280	285
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Met Asp Tyr Lys Val Ser 1 5		
aga agt ggg gag ata gta gaa gga gaa gta gaa gat tca gaa aag att 161		
Arg Ser Gly Ile Val Glu Gly Glu Val Glu Asp Ser Glu Lys Ile		
10 15 20		
gat tta cca cct ggt ttc aga ttt cac cca act gat gaa gaa ctt ata 209		
Asp Leu Pro Pro Gly Phe Arg Phe His Pro Thr Asp Glu Leu Ile		
25 30 35		
aca cac tat cta aga cca aag gtt gta aac tct ttt ttc tct gct ata 257		
Thr His Tyr Leu Arg Pro Lys Val Val Asn Ser Phe Phe Ser Ala Ile		
40 45 50		
gct att ggt gaa gtt gat ctc aac aaa gtc gag cct tgg gac ttg cct 305		
Ala Ile Gly Glu Val Asp Leu Asn Lys Val Glu Pro Trp Asp Leu Pro		

MBI15 Sequence Listing.ST25

55	60	65	70	
tgg aag gct aag ctt	ggg gaa aaa gag	tgg tac ttc ttt tgc	gta aga	353
Trp Lys Ala Lys Leu	Gly Glu Lys Trp	Tyr Phe Phe Cys Val	Arg	
75	80	85		
gac cga aaa tac ccg act	ggt tta aga acg aat	cgt gct act aaa gcc		401
Asp Arg Lys Tyr Pro Thr	Gly Leu Arg Thr Asn Arg	Ala Thr Lys Ala		
90	95	100		
ggt tat tgg aaa gct aca	ggg aaa gat aaa gag	atc ttc aaa ggg aaa		449
Gly Tyr Trp Lys Ala Thr	Gly Lys Asp Lys Glu Ile	Phe Lys Gly Lys		
105	110	115		
tct ctt gtt ggt atg aag	aaa aca ttg gtt ttc	tac aaa gga aga gct		497
Ser Leu Val Gly Met Lys	Thr Leu Val Phe Tyr Lys Gly Arg Ala			
120	125	130		
cct aaa gga gta aaa aca	aat tgg gtc atg cat	gag tat cga tta gaa		545
Pro Lys Gly Val Lys Thr	Asn Trp Val Met His	Glu Tyr Arg Leu Glu		
135	140	145	150	
ggc aaa ttc gct atc	gat aat ctc tct aaa acc	gct aag aac gaa tgt		593
Gly Lys Phe Ala Ile Asp Asn Leu Ser	Lys Thr Ala Lys Asn Glu Cys			
155	160	165		
gtt att agt cgt gtt ttt	cat aca ccg act gat	ggt acg aag gag cat		641
Val Ile Ser Arg Val Phe His	Thr Arg Thr Asp Gly Thr Lys Glu His			
170	175	180		
atg tcc gtt ggt tta cct	ccg ctg atg gat	tct cca tat cta aag		689
Met Ser Val Gly Leu Pro	Pro Leu Met Asp Ser Ser	Pro Tyr Leu Lys		
185	190	195		
agt aga gga caa gac	tct tta gcc ggg acc acc	ctt ggt ggg ttg ttg		737
Ser Arg Gly Gln Asp Ser	Leu Ala Gly Thr Thr	Leu Gly Gly Leu Leu		
200	205	210		
tct cac gtt acc tac	ttc tcc gac caa aca acc	gat gac aag agt ctt		785
Ser His Val Thr Tyr	Phe Ser Asp Gln Thr	Thr Asp Asp Lys Ser Leu		
215	220	225	230	
gtg gcc gat ttt aaa act	acc atg ttt ggt tcc gga tcg	act aac ttt		833
Val Ala Asp Phe Lys	Thr Thr Met Phe Gly Ser	Gly Ser Thr Asn Phe		
235	240	245		
tta cca aac ata ggt	tct cta cta gac ttc	gat cct ctg ttt cta caa		881
Leu Pro Asn Ile Gly Ser	Leu Asp Phe Asp Pro	Leu Phe Leu Gln		
250	255	260		
aac aat tct tca gta	ctg aag atg ttg ctt	gac aat gaa gaa acc caa		929
Asn Asn Ser Ser Val	Leu Lys Met Leu Leu Asp	Asn Glu Glu Thr Gln		
265	270	275		
ttt aag aag aat ctt	cac aat tca ggt	tca tca gag agt gaa cta aca		977
Phe Lys Asn Leu His	Asn Ser Gly Ser Ser	Glu Ser Glu Leu Thr		
280	285	290		
gcg agt tct tgg caa	ggt cac aat tct tat	ggt tcc act ggt cca gtg		1025
Ala Ser Ser Trp Gln	Gly His Asn Ser Tyr	Gly Ser Thr Gly Pro Val		
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Asn Leu Asp Cys Val	Trp Lys Phe			
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35 40 45

Ser Phe Phe Ser Ala Ile Ala Ile Gly Glu Val Asp Leu Asn Lys Val  
50 55 60

Glu Pro Trp Asp Leu Pro Trp Lys Ala Lys Leu Gly Glu Lys Glu Trp  
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Tyr Phe Phe Cys Val Arg Asp Arg Lys Tyr Pro Thr Gly Leu Arg Thr  
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Asn Arg Ala Thr Lys Ala Gly Tyr Trp Lys Ala Thr Gly Lys Asp Lys  
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Glu Ile Phe Lys Gly Lys Ser Leu Val Gly Met Lys Lys Thr Leu Val  
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Phe Tyr Lys Gly Arg Ala Pro Lys Gly Val Lys Thr Asn Trp Val Met  
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Ser Ser Pro Tyr Leu Lys Ser Arg Gly Gln Asp Ser Leu Ala Gly Thr  
195 200 205

Thr Leu Gly Gly Leu Leu Ser His Val Thr Tyr Phe Ser Asp Gln Thr  
210 215 220

Thr Asp Asp Lys Ser Leu Val Ala Asp Phe Lys Thr Thr Met Phe Gly  
225 230 235 240

Ser Gly Ser Thr Asn Phe Leu Pro Asn Ile Gly Ser Leu Leu Asp Phe  
245 250 255

Asp Pro Leu Phe Leu Gln Asn Asn Ser Ser Val Leu Lys Met Leu Leu  
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Asp Asn Glu Glu Thr Gln Phe Lys Lys Asn Leu His Asn Ser Gly Ser  
275 280 285

## MBI15 Sequence Listing.ST25

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MBI15 Sequence Listing.ST25

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Lys	Leu	Arg	Phe	Asp	Phe	Pro	Glu	Lys	Pro						
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Arg	Ser	Lys	Arg	Ser	Arg	Ser	Glu	Phe	Asp	Arg	Gln	Ser	Leu	Thr	Glu
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Val	Lys	Ser	His	Val	Cys	Ser	Ile	Cys	His	Lys	Ser	Phe	Ala	Thr	Gly
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## MBI15 Sequence Listing .ST25

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	gct tca gct tct aaa gtt gta gag aag aaa tgg tta gtg aaa gat gag	150
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	Lys Arg Asn Met Leu Gln Asp Glu Ile Asn Arg Val Asn Ser Glu Asn	
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	aag aag cta acc gaa atg tta gca aga gtc tgt gag aag tac tat gct	246
	Lys Lys Leu Thr Glu Met Leu Ala Arg Val Cys Glu Lys Tyr Tyr Ala	
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	75                      80                      85	
	gtt aac ttt cag aac aaa cag cta acg ggg aaa cga aaa caa gaa ctt	342
	Val Asn Phe Gln Asn Lys Gln Leu Thr Gly Lys Arg Lys Gln Glu Leu	
	90                      95                      100	
	gat gag ttt gtt agc tcc cca att gga ctc agt ctc gga cca atc gag	390
	Asp Glu Phe Val Ser Ser Pro Ile Gly Leu Ser Leu Gly Pro Ile Glu	
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	aac atc acc aac gat aaa gcg acg gtt tca acc gct tac ttt gct gct	438
	Asn Ile Thr Asn Asp Lys Ala Thr Val Ser Thr Ala Tyr Phe Ala Ala	
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	Glu Lys Ser Asp Thr Ser Leu Thr Val Lys Asp Gly Tyr Gln Trp Arg	
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	Lys Tyr Gly Gln Lys Ile Thr Arg Asp Asn Pro Ser Pro Arg Ala Tyr	
	155                      160                      165	
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	Phe Arg Cys Ser Phe Ser Pro Ser Cys Leu Val Lys Lys Val Gln	
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MBI15 Sequence Listing .ST25

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ggg acg att caa gag gtt ttg gtg caa caa atg gct tct tcg ttg acc															774
Gly	Thr	Ile	Gln	Glu	Val	Leu	Gln	Gln	Met	Ala	Ser	Ser	Leu	Thr	
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aaa gat cct aag ttc act gca gct ctt gcg act gct att tcc ggg aga															822
Lys	Asp	Pro	Lys	Phe	Thr	Ala	Ala	Leu	Ala	Thr	Ala	Ile	Ser	Gly	Arg
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Leu	Ile	Glu	Mis	Ser	Arg	Thr									
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Arg Val Cys Glu Lys Tyr Tyr Ala Leu Asn Asn Leu Met Glu Glu Leu															
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Gln Ser Arg Lys Ser Pro Glu Ser Val Asn Phe Gln Asn Lys Gln Leu															
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## MBI15 Sequence Listing.ST25

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Leu Val Ala Thr Tyr Glu Gly Thr His Asn His Thr Gly Pro His Ala  
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Ser Val Ser Arg Thr Val Lys Leu Asp Leu Val Gln Gly Gly Leu Glu  
 210 215 220

Pro Val Glu Glu Lys Lys Glu Arg Gly Thr Ile Gln Glu Val Leu Val  
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Gln Gln Met Ala Ser Ser Leu Thr Lys Asp Pro Lys Phe Thr Ala Ala  
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Thr Gln Leu Tyr Ser Leu Val Asp Ile Ala Arg Gly Ala Asn Arg Ile  
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Val Asn Tyr Lys Gly Lys Ser Tyr Ser Leu Lys Gly Cys Ile Ser Asp  
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Ala Lys Ser Met Arg Ser Leu Leu Val Gln Gln Met Gly Phe Pro Ile  
115 120 125

gac tct att ctc atg ctc aca gaa gat gaa gcc agc ccg cag aga ata 432  
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130 135 140

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MBI15 Sequence Listing .ST25

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Gln Gln Asn Asp Tyr Asn Gly Asp Glu Ile Asp Gly Gln Asp Glu Ala			
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ttg tgc cct tta gac cat gaa aca gaa gga aaa atc att gat gac gag			624
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195	200	205	
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Ser Val Arg Ala Tyr Lys Gly Thr Asp Gly Gly Ala Ala Phe Cys Phe			
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Ser Ala Cys Asp Asp Asp Glu Ser Ser Gly Tyr Thr Pro Val Phe Thr			
275	280	285	
ggg aag aac aca gga gcc atg act tat agc ttc ata aag gcg gtg aag			912
Gly Lys Asn Thr Gly Ala Met Thr Tyr Ser Phe Ile Lys Ala Val Lys			
290	295	300	
aca gct gga cca gca ccc acg tat ggc cac ctg ctt aac ctt atg tgt			960
Thr Ala Gly Pro Ala Pro Thr Tyr Gly His Leu Leu Asn Leu Met Cys			
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Ser Ala Ile Arg Glu Ala Gln Ser Arg Leu Ala Phe Asn Gly Asp Tyr			
325	330	335	
aca agc tct gat gca tcc gcg gag cca ctg cta aca tca tct gag gaa			1056
Thr Ser Ser Asp Ala Ser Ala Glu Pro Leu Leu Thr Ser Ser Glu Glu			
340	345	350	
ttt gac gtg tac gcg aca aag ttt gta ctc tga atgctgtaca tatgatgctg			1109
Phe Asp Val Tyr Ala Thr Lys Phe Val Leu			
355	360		
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## MBI15 Sequence Listing.ST25

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Ile His Gly Phe Gln Gln Leu Leu Arg Gln His Gln Pro Gln His His  
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Glu Gln Gln Gln Gln Met Met Ala Gln Pro Pro Pro Arg Leu Leu  
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Glu Pro Leu Pro Ser Pro Phe Gly Lys Lys Arg Ala Val Leu Cys Gly  
85 90 95

Val Asn Tyr Lys Gly Lys Ser Tyr Ser Leu Lys Gly Cys Ile Ser Asp  
100 105 110

Ala Lys Ser Met Arg Ser Leu Leu Val Gln Gln Met Gly Phe Pro Ile  
115 120 125

Asp Ser Ile Leu Met Leu Thr Glu Asp Glu Ala Ser Pro Gln Arg Ile  
130 135 140

Pro Thr Lys Arg Asn Ile Arg Lys Ala Met Arg Trp Leu Val Glu Gly  
145 150 155 160

Asn Arg Ala Arg Asp Ser Leu Val Phe His Phe Ser Gly His Gly Ser  
165 170 175

Gln Gln Asn Asp Tyr Asn Gly Asp Glu Ile Asp Gly Gln Asp Glu Ala  
180 185 190

Leu Cys Pro Leu Asp His Glu Thr Glu Gly Lys Ile Ile Asp Asp Glu  
195 200 205

Ile Asn Arg Ile Leu Val Arg Pro Leu Val His Gly Ala Lys Leu His  
210 215 220

Ala Val Ile Asp Ala Cys Asn Ser Gly Thr Val Leu Asp Leu Pro Phe  
225 230 235 240

Ile Cys Arg Met Glu Arg Asn Gly Ser Tyr Glu Trp Glu Asp His Arg  
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Ser Val Arg Ala Tyr Lys Gly Thr Asp Gly Gly Ala Ala Phe Cys Phe  
260 265 270

Ser Ala Cys Asp Asp Asp Glu Ser Ser Gly Tyr Thr Pro Val Phe Thr  
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Gly Lys Asn Thr Gly Ala Met Thr Tyr Ser Phe Ile Lys Ala Val Lys  
290 295 300

Thr Ala Gly Pro Ala Pro Thr Tyr Gly His Leu Leu Asn Leu Met Cys  
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Ser Ala Ile Arg Glu Ala Gln Ser Arg Leu Ala Phe Asn Gly Asp Tyr

## MBI15 Sequence Listing.ST25

325

330

335

Thr Ser Ser Asp Ala Ser Ala Glu Pro Leu Leu Thr Ser Ser Glu Glu  
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Phe Asp Val Tyr Ala Thr Lys Phe Val Leu  
355 360

## INTERNATIONAL SEARCH REPORT

International application No.

PCT/US00/31418

**A. CLASSIFICATION OF SUBJECT MATTER**

IPC(7) : A01H 1/00, 5/00; C12N 5/14, 15/82  
 US CL : 435/320.1, 419, 468; 800/278, 279, 287, 301, 305-310, 312, 314, 317, 320, 322

According to International Patent Classification (IPC) or to both national classification and IPC

**B. FIELDS SEARCHED**

Minimum documentation searched (classification system followed by classification symbols)

U.S. : 435/320.1, 419, 468; 800/278, 279, 287, 301, 305-310, 312, 314, 317, 320, 322

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practicable, search terms used)  
EAST, USPAT, STN, Agricola, CaPlus, Biosis, Embase**C. DOCUMENTS CONSIDERED TO BE RELEVANT**

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	WO 97/47183 A1 (PURDUE RESEARCH FOUNDATION) 18 December 1997 (18.12.1997), entire reference.	1-9, 12, 13, 25
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Y		10, 11, 26, 27
X	US 5,939,601 (KLESSIG et al) 17 August 1999 (17.08.1999), entire reference.	1-9, 12, 13, 25
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Y		10, 11, 26, 27
A	Database Genbank on NCBI, US National Library of Medicine, (Bethesda, MD, USA) No. AB009055, SATO, S. et al 'Structural analysis of Arabidopsis thaliana chromosome 5. IV. Sequence features of the regions of 1,456,315 bp covered by nineteen physically assigned P1 and TAC clones. 27 December 2000, DNA RES. 1998, Vol. 5, No. 1, pages 41-54, see bases 16,003-16,490, 16,571-16,683 and 16,780-17,365.	1-13, 25-27

 Further documents are listed in the continuation of Box C.

See patent family annex.

• Special categories of cited documents:	"T"	later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention
"A" document defining the general state of the art which is not considered to be of particular relevance	"X"	document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone
"E" earlier application or patent published on or after the international filing date	"Y"	document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art
"L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)	"&"	document member of the same patent family
"O" document referring to an oral disclosure, use, exhibition or other means		
"P" document published prior to the international filing date but later than the priority date claimed		

Date of the actual completion of the international search

23 February 2001 (23.02.2001)

Date of mailing of the international search report

09 MAR 2001

Name and mailing address of the ISA/US  
 Commissioner of Patents and Trademarks  
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**INTERNATIONAL SEARCH REPORT**

International application No.

PCT/US00/31418

**Box I Observations where certain claims were found unsearchable (Continuation of Item 1 of first sheet)**

This international report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:

1.  Claim Nos.:  
because they relate to subject matter not required to be searched by this Authority, namely:
  
2.  Claim Nos.:  
because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically:
  
3.  Claim Nos.: 14  
because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).

**Box II Observations where unity of invention is lacking (Continuation of Item 2 of first sheet)**

This International Searching Authority found multiple inventions in this international application, as follows:  
Please See Continuation Sheet

1.  As all required additional search fees were timely paid by the applicant, this international search report covers all searchable claims.
2.  As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3.  As only some of the required additional search fees were timely paid by the applicant, this international search report covers only those claims for which fees were paid, specifically claims Nos.:
  
4.  No required additional search fees were timely paid by the applicant. Consequently, this international search report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.: 1-13 & 25-27 and SEQ ID NOS 1&2

**Remark on Protest**  

The additional search fees were accompanied by the applicant's protest.

No protest accompanied the payment of additional search fees.

**INTERNATIONAL SEARCH REPORT**

Int'l. application No.

PCT/US00/31418

**BOX II. OBSERVATIONS WHERE UNITY OF INVENTION IS LACKING** This application contains the following inventions or groups of inventions which are not so linked as to form a single general inventive concept under PCT Rule 13.1. In order for all inventions to be examined, the appropriate additional examination fees must be paid.

Groups I-XXIX, claim(s) 1-14 and 25-27, drawn to a transgenic plant having modified seed characteristics, polynucleotides and vectors for producing said transgenic plant and a method of making said transgenic plant. Applicant must elect one pair of sequences (one nucleic acid and the corresponding amino acid translation) to be examined, *i.e.* SEQ ID NO: 1 and 2 in Group I, SEQ ID NO: 3 and 4 in Group II, SEQ ID NO: 5 and 6 in Group III, etc.

Group XXX, claim(s) 15-17, drawn to a method of identifying a factor that is modulated.

Group XXXI, claims(s) 18, drawn to a method of identifying a molecule that modulates activity or expression of a polynucleotide or polypeptide.

Group XXXII, claims(s) 19 and 20, drawn to an integrated computer system.

Group XXXIII, claim(s) 21-24, drawn to a method for identifying a polynucleotide sequence comprising selecting a nucleic acid sequence from a database that meets a selected sequence criteria.

The inventions listed as Groups I-XXXIII do not relate to a single general inventive concept under PCT Rule 13.1 because, under PCT Rule 13.2, they lack the same or corresponding special technical features for the following reasons:

The inventions listed as Groups I-XXXIII do not relate to a single general inventive concept under PCT Rule 13.1 because, under PCT Rule 13.2, they lack the same or corresponding special technical features for the following reasons: Groups I-XXIX are drawn to a transgenic plant and a method of producing said plant with a nucleic acid sequence. The methods of Groups I-XXIX differ from each other in that they are directed to a plant transformation method and transgenic plant with a structurally and functionally distinct nucleic acid sequence which encodes a structurally and functionally distinct amino acid sequence. In addition, Groups XXX, XXXI and XXXIII are different methods from any of Groups I-XXIX in that they have different method steps and different end products, and Group XXXII requires a computer system. Thus, there is no single special technical feature, which links the inventions of Groups I-XXXIII under PCT Rule 13.2.